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L41

(FILE 'HOME' ENTERED AT 15:22:31 ON 02 FEB 2004) SET COST OFF FILE 'HCAPLUS' ENTERED AT 15:22:50 ON 02 FEB 2004 E ALBUMIN/CT 753 S E3 L1L2 132 S E11 E E47+ALL 80101 S E2+NT L3 E E33+ALL 566 S E3, E2 L4L5 25218 S E2+NT L6 157881 S ?ALBUMIN? L7 181833 S L1-L6  $\Gamma8$ 2969 S BDNF OR BD NF 2881 S BRAIN DERIVED NEUROTROPHIC FACTOR L9 2883 S (BD OR BRAIN DERIVED) () (NF OR NEUROTROPHIC FACTOR) L10 E NEUROTROPHIC FACTOR/CT 141 S E10 L11L12 2554 S E26 E E25+ALL 789 S E3-E5 AND BRAIN DERIVED L13 L14 679 S E12,E13 L15 3242 S E2+NT (L) BRAIN DERIVED L16 64 S L7 AND L8-L15 L17 19234 S INTERFERONALPHA OR ALPHAINTERFERON OR INTERFERONBETA OR BETAI E INTERFERON/CT 302 S E3-E19 L18 18390 S E85-E101 E INTERFERONS/CT E E3+ALL 18391 S E7, E6 (L) (ALPHA OR BETA) L20. 546 S L7 AND L17-L20 L21 L22 2340 S TIMP()(I OR 1) FILE 'REGISTRY' ENTERED AT 15:29:36 ON 02 FEB 2004 1 S 140208-24-8 L23 FILE 'HCAPLUS' ENTERED AT 15:30:37 ON 02 FEB 2004 2026 S L23 L24 L25 859 S TISSUE INHIBITOR (1W) METALLOPROTEINASE 1 27 S METALLOPROTEINASE INHIBITOR 1 L26 L27 651 S TIMP1 12 S FIBROBLAST COLLAGENASE INHIBITOR L28 L29 91 S L7 AND L22, L24-L28 L30 678 S L16, L21, L29 L31 9815 S IFNALPHA OR IFNBETA OR ALPHAIFN OR BETAIFN OR IFN(A) (ALPHA OR L32 119 S L7 AND L31 L33 700 S L30, L32 L34 62 S L33 AND (FUSION OR FUSE OR FUSED OR FUSES OR FUSING) L35 167 S L33 AND RECOMBIN? 44 S L33 AND CHIMER? L36 L37 202 S L34-L36 E ROSEN C/AU L38 27 S E3, E4 E ROSEN CRAIG/AU 625 S E3-E5 L39 E HASELTINE W/AU 302 S E3, E4, E7-E10 L40

10 S L33 AND L38-L40

E HUMAN GENOME SCI/PA, CS

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975 S E5-E37
             13 S L33 AND L42
L43
             13 S L41, L43
L44
L45
             13 S L44 AND L37
              9 S L45 AND (SHELFLIFE OR SHELF LIFE)
L46
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L47
                SEL DN AN 1 4
              2 S L47 NOT E1-E6
L48
             11 S L46, L48
L49
                SEL RN
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L50
          11933 S E9
                E E9+ALL
           3795 S E3, E4
L51
L52
              5 S L51 AND L33
             29 S L50 AND L33
L53
L54
             34 S L49, L52, L53
             27 S L54 AND ALBUMIN
L55
              7 S L54 NOT L55
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L57
            159 S L37 AND ALBUMIN
L58
            132 S L57 NOT L43-L49, L52-L56
              6 S L58 AND L16
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L60
              7 S L58 AND L29
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             96 S L61 AND (PD<=20000412 OR PRD<=20000412 OR AD<=20000412)
L62
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             18 S L62 AND E1-E54
L63
L64
             29 S L49, L63 AND L1-L22, L24-L63
L65
             29 S L64 AND ?ALBUMIN?
             29 S L64 AND (INF? OR INTERFERON OR TIMP? OR NEUROTROPHIC?)
L66
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ن رئيس :

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FILE 'HCAPLUS' ENTERED AT 16:00:16 ON 02 FEB 2004
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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FILE COVERS 1907 - 2 Feb 2004 VOL 140 ISS 6 FILE LAST UPDATED: 1 Feb 2004 (20040201/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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L66 ANSWER 1 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN AN 2003:571103 HCAPLUS
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DN 139:122690

ED Entered STN: 25 Jul 2003

TI Albumin fusion proteins for prolonged shelf-life of therapeutic proteins

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ΙN
     Ballance, David James; Turner, Andrew John; Rosen, Craig A.; Haseltine,
     William A.
PA
     Human Genome Sciences, Inc., USA; Delta Biotechnology Limited; Principia
     Pharmaceutical Corporation
     PCT Int. Appl., 598 pp.
SO
     CODEN: PIXXD2
DT
     Patent
     English
LA
IC
     ICM C12N
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3
FAN.CNT 2
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     PATENT NO.
                      KIND
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                      A2
                            20030724
                                           WO 2002-US40891 20021223
     WO 2003060071
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
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                       Ρ
                            20021023
     US 2002-423623P
                       Ρ
                            20021105
     The present invention encompasses albumin fusion proteins. Many
AΒ
     therapeutic proteins in their native state or when recombinantly produced .
     are typically labile mols. exhibiting short shelf-lives, particularly when
     formulated in aqueous solns.; fusions of the therapeutic protein with human
     serum albumin have a longer serum half-life and/or stabilized activity in
     solution (or in a pharmaceutical composition) in vitro and/or in vivo than the
     corresponding unfused therapeutic mols. Thus, albumin fusion proteins are
     provided comprising granulocyte colony-stimulating factor, interleukin 2,
     parathormone, erythropoietin, interferon \beta, interferon \alpha2,
     interferon A/D hybrid, a single-chain insulin analog, growth hormone, and
     (7-36) GLP-1. Nucleic acid mols. encoding the albumin fusion proteins of
     the invention are also encompassed by the invention, as are vectors containing
     these nucleic acids, host cells transformed with these nucleic acids
     vectors, and methods of making the albumin fusion proteins of the
     invention and using these nucleic acids, vectors, and/or host cells.
     Addnl. the present invention encompasses pharmaceutical compns. comprising
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albumin fusion proteins and methods of treating or preventing diseases,

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disorders or conditions related to diabetes mellitus using albumin fusion proteins of the invention. albumin fusion therapeutic protein shelflife Animal cell line (293, recombinant expression host; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Animal cell line (CHO, recombinant expression host; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Animal cell line (NSO, recombinant expression host; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (antiviral, T1249 peptide inhibitor derived from HIV-1; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Antidiabetic agents Human Linking agents Molecular cloning (human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Fusion proteins (chimeric proteins) Interleukin 2 RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Signal peptides RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Animal cell (mammalian, recombinant expression host; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Diabetes mellitus (non-insulin-dependent, treatment of; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Protein sequences (of human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Plasmid vectors (pC4; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Plasmid vectors (pEE12.1; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Plasmid vectors (pSAC35; human serum albumin fusion proteins for prolonged shelf-life of therapeutic proteins) Saccharomyces cerevisiae Yeast (recombinant expression host that is glycosylation and protease-deficient; human serum albumin fusion proteins for prolonged

Albumins, biological studies IT RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

shelf-life of therapeutic proteins)

- 75

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(serum; human serum albumin fusion proteins for prolonged shelf-life of
        therapeutic proteins)
IT
     Interferons
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha 2; human serum albumin fusion proteins for prolonged shelf-life
        of therapeutic proteins)
ΙT
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha; human serum albumin fusion proteins for prolonged shelf-life
        of therapeutic proteins)
ΙT
     Interferons
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha AD; human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
ΙT
     Interferons
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (β; human serum albumin fusion proteins for prolonged shelf-life
        of therapeutic proteins)
ΙT
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                    562119-83-9P
                                    562119-85-1DP, Albumin (human),
     subfragments, fusion products
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (amino acid sequence; human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
                                 9004-10-8P, Insulin, biological studies
ΙT
     9002-64-6P, Parathormone
     11096-26-7P, Erythropoietin
                                   89750-14-1P, Glucagon-like peptide I
     143011-72-7P, Granulocyte colony-stimulating factor
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (human serum albumin fusion proteins for prolonged shelf-life of
        therapeutic proteins)
ΙT
     562119-84-0
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     study); USES (Uses)
        (nucleotide sequence; human serum albumin fusion proteins for prolonged
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562131-47-9

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   shelf-life of therapeutic proteins)
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        (unclaimed nucleotide sequence; albumin fusion proteins for prolonged
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     RL: PRP (Properties)
        (unclaimed protein sequence; albumin fusion proteins for prolonged
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shelf-life of therapeutic proteins)

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        (unclaimed sequence; albumin fusion proteins for prolonged shelf-life
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DN
     139:122689.
     Entered STN: 25 Jul 2003
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     Albumin fusion proteins for prolonged shelf-
ΤI
     life of therapeutic proteins
ΙN
     Rosen, Craig A.; Haseltine, William A.
PΑ
     Human Genome Sciences, Inc., USA
SO
     PCT Int. Appl., 1086 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
IC
     ICM C07K
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3
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                      KIND DATE
                                            APPLICATION NO.
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و و السيتون

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US 2002-420246P
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    US 2002-423623P
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    The present invention encompasses albumin fusion
AB
    proteins. Many therapeutic proteins in their native state or when
    recombinantly produced are typically labile mols. exhibiting short
    shelf-lives, particularly when formulated in aqueous solns.;
    fusions of the therapeutic protein with human serum
    albumin have a longer serum half-life and/or stabilized activity
    in solution (or in a pharmaceutical composition) in vitro and/or in vivo than
the
    corresponding unfused therapeutic mols. Thus, albumin
    fusion proteins are provided comprising interferon .
    beta., interferon \alpha 2, insulin, bone
    morphogenetic protein 9, glucagon-like peptide-I(7-36), a hybrid
    interferon A/D, and extendin 4. Nucleic acid mols. encoding the
    albumin fusion proteins of the invention are also
     encompassed by the invention, as are vectors containing these nucleic acids,
    host cells transformed with these nucleic acids vectors, and methods of
    making the albumin fusion proteins of the invention
     and using these nucleic acids, vectors, and/or host cells. Addnl. the
    present invention encompasses pharmaceutical compns. comprising
    albumin fusion proteins and methods of treating or
    preventing diseases, disorders or conditions related to diabetes mellitus
     using albumin fusion proteins of the invention.
ST
    albumin fusion therapeutic protein shelflife
ΙT
    Animal cell line
        (293, recombinant expression host; human serum
        albumin fusion proteins for prolonged shelf
        -life of therapeutic proteins)
    Animal cell line
IT
        (CHO, recombinant expression host; human serum
        albumin fusion proteins for prolonged shelf
        -life of therapeutic proteins)
    Animal cell line
IT
        (NSO, recombinant expression host; human serum
        albumin fusion proteins for prolonged shelf
        -life of therapeutic proteins)
IT
    Metabolism, animal
        (disorder, treatment of; human serum albumin fusion
        proteins for prolonged shelf-life of therapeutic
        proteins)
ΤТ
    Antidiabetic agents
    Antiobesity agents
    Cardiovascular agents
    Human
    Linking agents
    Molecular cloning
        (human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
ΙT
     Fusion proteins (chimeric proteins)
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
ΙT
     Signal peptides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
     Diabetes mellitus
ΙT
        (insulin-dependent, treatment of; human serum albumin
```

fusion proteins for prolonged shelf-life of

```
therapeutic proteins)
ΙT
    Animal cell
        (mammalian, recombinant expression host; human serum
        albumin fusion proteins for prolonged shelf
        -life of therapeutic proteins)
ΙT
     Nerve, disease
        (neuropathy, treatment of; human serum albumin fusion
        proteins for prolonged shelf-life of therapeutic
        proteins)
ΙT
     Diabetes mellitus
        (non-insulin-dependent, treatment of; human serum albumin
        fusion proteins for prolonged shelf-life of
        therapeutic proteins)
     Protein sequences
ΙT
        (of human serum albumin fusion proteins for
        prolonged shelf-life of therapeutic proteins)
     Plasmid vectors
TT
        (pC4; human serum albumin fusion proteins for
        prolonged shelf-life of therapeutic proteins)
ΙT
     Plasmid vectors
        (pEE12.1; human serum albumin fusion proteins for
        prolonged shelf-life of therapeutic proteins)
ΙT
     Plasmid vectors
        (pSAC35; human serum albumin fusion proteins for
        prolonged shelf-life of therapeutic proteins) .
ΙΤ
     Saccharomyces cerevisiae
     Yeast
        (recombinant expression host that is glycosylation and
        protease-deficient; human serum albumin fusion
        proteins for prolonged shelf-life of therapeutic
        proteins)
IT
    Eye, disease
        (retinopathy, treatment of; human serum albumin
        fusion proteins for prolonged shelf-life of
        therapeutic proteins)
     Albumins, biological studies
IT
    RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (serum; human serum albumin fusion proteins for
        prolonged shelf-life of therapeutic proteins)
     Cardiovascular system, disease
ΙT
     Endocrine system, disease
     Heart, disease
     Hyperglycemia
    Kidney, disease
     Nervous system, disease
     Obesity
        (treatment of; human serum albumin fusion proteins
        for prolonged shelf-life of therapeutic proteins)
ΙT
     Interferons
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha 2; human serum albumin
        fusion proteins for prolonged shelf-life of
        therapeutic proteins)
TT
     Interferons
     RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha ; human serum albumin fusion
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proteins for prolonged shelf-life of therapeutic

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proteins)
TT
    Interferons
    RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\alpha AD; human serum albumin
        fusion proteins for prolonged shelf-life of
        therapeutic proteins)
ΙT
    Interferons
    RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (\beta ; human serum albumin fusion
        proteins for prolonged shelf-life of therapeutic
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TΤ
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        (Unclaimed; albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
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    561347-54-4DP, Albumin (human), subfragments, fusion
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     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (amino acid sequence; human serum albumin fusion
        proteins for prolonged shelf-life of therapeutic
       proteins)
ΙT
     9004-10-8P, Insulin, biological studies
                                               107444-51-9P,
     (7-36)Glucagon-like peptide 1 amide
                                          141732-76-5P, Extendin 4
    305835-60-3P, Bone morphogenetic protein 9
    RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); USES (Uses)
        (human serum albumin fusion proteins for prolonged
        shelf-life of therapeutic proteins)
TΤ
    50-99-7, D-Glucose, biological studies
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (maintenance of basel level of; human serum albumin
        fusion proteins for prolonged shelf-life of
        therapeutic proteins)
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    study); USES (Uses)
        (nucleotide sequence; human serum albumin fusion
       .proteins for prolonged shelf-life of therapeutic
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        proteins for prolonged shelf-life of therapeutic
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     561354-79-8
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                                                  561354-82-3
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                    561354-96-9
                                   561354-97-0
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
        proteins for prolonged shelf-life of therapeutic
        proteins)
ΙT
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                                   561350-51-4
                                                                 561350-53-6
                                                  561350-52-5
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                                                           561352-68-9
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                                            561353-80-8
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               561353-89-7
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                                                           561353-92-2
561353-93-3
               561353-94-4
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                                            561353-96-6
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RL: PRP (Properties)
   (unclaimed protein sequence; albumin fusion
   proteins for prolonged shelf-life of therapeutic
   proteins)
                                             40958-31-4, Somatostatin (sheep
33017-11-7, Proinsulin C-peptide (human)
          82177-09-1
                         85482-68-4
                                       85734-71-0
                                                    122024-47-9
reduced)
                             131748-18-0
                                            131748-19-1
                                                           157654-59-6
125677-89-6
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               170098-75-6
166980-40-1
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               477953-25-6
                             477953-26-7
                                            477953-27-8
                                                           477953-28-9
367273-48-1
               477.953-30-3
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                                            477953-32-5
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477953-29-0
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                             478188-11-3
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                                                           561304-79-8
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561304-80-1
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                             561304-83-4
                                            561304-84-5
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               561304-87-8
                             561304-88-9
                                            561304-92-5
                                                           561304-95-8
561304-86-7
RL: PRP (Properties)
   (unclaimed sequence; albumin fusion proteins for
   prolonged shelf-life of therapeutic proteins)
ANSWER 3 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
2003:300832 HCAPLUS
138:326508
Entered STN:
              18 Apr 2003
Albumin fusion proteins with therapeutic proteins for
improved shelf-life
Rosen, Craig A.; Haseltine, William A.
Human Genome Sciences, Inc., USA
PCT Int. Appl., 457 pp.
CODEN: PIXXD2
Patent
English
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-7.

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CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3, 15
FAN.CNT 1
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
     PATENT NO.
                           _____
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                                           -----
                      A2
                                           WO 2002-US31794
                                                            20021004
PΙ
     WO 2003030821
                            20030417
                      A3
                            20031211
     WO 2003030821
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD,
             RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
                            20011005
PRAI US 2001-327281P
                      Ρ
     The present invention encompasses fusion proteins of
     albumin with various therapeutic proteins. Therapeutic proteins
     may be stabilized to extend the shelf-life, and/or to
     retain the therapeutic protein's activity for extended periods of time in
     solution, in vitro and/or in vivo, by genetically or chemical fusing
     or conjugating the therapeutic protein to albumin or a fragment
    or variant of albumin. Use of albumin fusion
     proteins may also reduce the need to formulate the protein solns. with
     large excesses of carrier proteins to prevent loss of therapeutic proteins
     due to factors such as binding to the container. Nucleic acid mols.
     encoding the albumin fusion proteins of the invention
     are also encompassed by the invention, as are vectors containing these nucleic
     acids, host cells transformed with these nucleic acids vectors, and
     methods of making the albumin fusion proteins of the
     invention and using these nucleic acids, vectors, and/or host cells.
     Thus, plasmid vectors are constructed in which DNA encoding the desired
     therapeutic protein may be inserted for expression of the albumin
     fusion proteins in yeast (pPPC0005) and mammalian cells (pC4:HSA).
     Yeast-derived signal sequences from Saccharomyces cerevisiae invertase
     SUC2 gene, or the stanniocalcin or native human serum albumin
     signal peptides, are used for secretion in yeast or mammalian systems,
           Thus, the fusion product of human growth hormone with
     residues 1-387 of human serum albumin retains essentially intact
     biol. activity after 5 wk of incubation in tissue culture media at
     37°, whereas recombinant human growth hormone used as
     control lost its biol. activity in the first week. Although the potency
     of the albumin fusion proteins is slightly lower than
     the unfused counterparts in rapid bioassays, their biol. stability results
     in much higher biol. activity in the longer term in vitro assay or in vivo
             Addnl., the present invention encompasses pharmaceutical compns.
     Comprising albumin fusion proteins and methods of
     treating, preventing, or ameliorating diseases, disorders or conditions
     using albumin fusion proteins of the invention.
ST
     albumin fusion therapeutic protein shelflife
IT
     Drug delivery systems
     Gene therapy
     Human
     Molecular cloning
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     Fusion proteins (chimeric proteins)
       Interferons
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
IT
     Signal peptides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     Peptides, biological studies
IT
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (linkers; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT.
     Animal cell
        (mammalian, recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Plasmid vectors
        (pScCHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
TT
     Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Linking agents
        (peptide; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Saccharomyces cerevisiae
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
TT
     Albumins, biological studies
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (serum; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (signal sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TT
     Antibodies
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (single chain; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (therapeutic; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Interferons
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(\alpha ; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     9002-72-6DP, Growth hormone, fusion proteins with
ΙT
     albumin
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     511566-72-6DP, Albumin (human blood serum), full-length or
ΙT
     subfragment fusion proteins
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (amino acid sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     511566-73-7
     RL: BUU (Biological use, unclassified); PRP (Properties); BIOL (Biological
     study); USES (Uses)
        (nucleotide sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     511603-12-6
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                                                511603-15-9
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     511603-17-1
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                                                              511603-21-7
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     511603-67-1
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     122024-47-9
                                 367273-46-9
                                                367273-47-0
                                                              367273-48-1
TΤ
                   131748-18-0
     RL: PRP (Properties)
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
    ANSWER 4 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
AN
     2003:125793 HCAPLUS
DN
     138:297265
ED
     Entered STN: 19 Feb 2003
ΤI
     An IFN-\beta -Albumin Fusion
     Protein That Displays Improved Pharmacokinetic and Pharmacodynamic
     Properties in Nonhuman Primates
ΑU
     Sung, Cynthia; Nardelli, Bernardetta; LaFleur, David W.; Blatter, Erich;
     Corcoran, Marta; Olsen, Henrik S.; Birse, Charles E.; Pickeral, Oxana K.;
     Zhang, Junli; Shah, Devanshi; Moody, Gordon; Gentz, Solange; Beebe, Lisa;
     Moore, Paul A.
CS
     Human Genome Sciences, Inc., Rockville, MD, 20850, USA
     Journal of Interferon and Cytokine Research (2003), 23(1), 25-36
SO
     CODEN: JICRFJ; ISSN: 1079-9907
PB
     Mary Ann Liebert, Inc.
DT
     Journal
LA
     English
CC
     1-7 (Pharmacology)
     Section cross-reference(s): 15
AΒ
     The long half-life and stability of human serum albumin (HSA)
     make it an attractive candidate for fusion to short-lived
     therapeutic proteins. Albuferon beta (Human Genome Sciences [HGS], Inc.,
     Rockville, MD) is a novel recombinant protein derived from a
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 $\beta$  ; IFN- $\beta$  -albumin

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gene fusion of interferon-\beta
                             (
IFN-β ) and HSA. In vitro, Albuferon beta displays
antiviral and antiproliferative activities and triggers the IFN-stimulated
response element (ISRE) signal transduction pathway. Array anal. of 5694
independent genes in Daudi-treated cells revealed that Albuferon beta and
IFN-\beta induce the expression of an identical set of
.30 genes, including 9 previously not identified.
                                                   In rhesus monkeys
administered a dose of 50 µg/kg i.v. or s.c. or 300 µg/kg s.c.,
Albuferon beta demonstrated favorable pharmacokinetic properties. S.c.
bioavailability was 87%, plasma clearance at 4.7-5.7 mL/h/kg was approx.
140-fold lower than that of IFN-\beta , and the
terminal half-life was 36-40 h compared with 8 h for IFN-.
        Importantly, Albuferon beta induced sustained increases in
serum neopterin levels and 2',5'-oligoadenylate synthetase (2',5'-OAS)
mRNA expression. At a molar dose equivalent to one-half the dose of
IFN-β , Albuferon beta elicited comparable neopterin
responses and significantly higher 2',5'-OAS mRNA levels in rhesus
monkeys. The enhanced in vivo pharmacol. properties of IFN-.
beta. when fused to serum albumin suggest a
clin. opportunity for improved IFN-\beta therapy.
interferon beta albumin fusion
protein albuferon beta pharmacokinetic pharmacodynamic
Fusion proteins (chimeric proteins)
RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT
(Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); PREP
(Preparation); USES (Uses)
   (IFN-\beta -HSA; IFN-\beta -
   albumin fusion protein with retained biol. activities
   and improved pharmacokinetic and pharmacodynamic properties of
   IFN-\beta in primates)
Antiviral agents
Human
Macaca mulatta
Pharmacodynamics
Pharmacokinetics
Signal transduction, biological
    (IFN-\beta -albumin fusion
   protein with retained biol. activities and improved pharmacokinetic and
   pharmacodynamic properties of IFN-\beta in
   primates)
Genetic element
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (ISRE (interferon-stimulated response element); IFN
   -\beta -albumin fusion protein with
   retained biol. activities and improved pharmacokinetic and
   pharmacodynamic properties of IFN-\beta in
   primates)
Transcriptional regulation
    (activation; IFN-\beta -albumin
   fusion protein with retained biol. activities and improved
   pharmacokinetic and pharmacodynamic properties of IFN-
   \beta in primates)
Cell proliferation
   (inhibition; IFN-\beta -albumin
   fusion protein with retained biol. activities and improved
   pharmacokinetic and pharmacodynamic properties of IFN-
      in primates)
Albumins, biological studies
RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT
(Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); PREP
(Preparation); USES (Uses)
    (serum, human, fusion protein with IFN-
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fusion protein with retained biol. activities and improved pharmacokinetic and pharmacodynamic properties of IFNin primates) Interferons RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (β , fusion protein with albumin; IFN- $\beta$  -albumin fusion protein with retained biol. activities and improved pharmacokinetic and pharmacodynamic properties of  $IFN-\beta$  in primates) 507485-69-0P, Albuferon beta RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (IFN- $\beta$  -HSA; IFN- $\beta$  albumin fusion protein with retained biol. activities and improved pharmacokinetic and pharmacodynamic properties of **IFN-** $\beta$  in primates) 69106-44-1, 2',5'-Oligoadenylate synthetase 2009-64-5, Neopterin RL: BSU (Biological study, unclassified); BIOL (Biological study) (IFN- $\beta$  -albumin fusion protein with retained biol. activities and improved pharmacokinetic and pharmacodynamic properties of IFN- $\beta$  in primates) RE.CNT THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Brumell, J; J Immunol 1999, V163, P3388 HCAPLUS (2) Chuang, V; Pharm Res 2002, V19, P569 (3) Durelli, L; Lancet 2002, V359, P1453 HCAPLUS (4) Eisen, M; Proc Natl Acad Sci USA 1998, V95, P14863 HCAPLUS (5) Fierlbeck, G; J Interferon Cytokine Res 1996, V16, P777 MEDLINE (6) Fine, H; Clin Cancer Res 1997, V3, P381 HCAPLUS (7) Fukutomo, T; J Hepatol 2001, V34, P100 (8) Glue, P; Clin Pharmacol Ther 2000, V68, P556 HCAPLUS (9) Grace, M; J Interferon Cytokine Res 2001, V21, P1103 HCAPLUS (10) Gutterman, J; Proc Natl Acad Sci USA 1994, V91, P1198 HCAPLUS (11) Imaizumi, T; J Leukocyte Biol 2002, V72, P486 HCAPLUS (12) Jacobs, L; N Engl J Med 2000, V343, P898 HCAPLUS (13) Karsan, A; Blood 1996, V87, P3089 HCAPLUS (14) Kho, C; J Biol Chem 1997, V272, P13426 HCAPLUS (15) Lafleur, D; J Biol Chem 2001, V276, P39765 HCAPLUS (16) Leaman, D; J Biol Chem 2002, V277, P28504 HCAPLUS (17) Lindsay, K; Hepatology 2001, V34, P395 HCAPLUS (18) Lukashok, S; J Virol 2000, V74, P4705 HCAPLUS (19) Maeyer, E; The Cytokine Handbook, 3rd ed 1998, P491 (20) Marques, J; Thromb Haemost 2001, V86, P902 HCAPLUS (21) Osborn, B; Eur J Pharmacol 2002, V456, P149 HCAPLUS (22) Osborn, B; J Pharmacol Exp Ther 2002, V303, P540 HCAPLUS (23) Paty, D; Neurology 1993, V43, P662 MEDLINE (24) Pellegrini, S; Mol Cell Biol 1989, V9, P4605 HCAPLUS (25) Pepinsky, R; J Pharmacol Exp Ther 2001, V297, P1059 HCAPLUS (26) Peters, T; All About Albumin 1996 (27) Pferrer, L; Cancer Res 1998, V58, P2489 (28) Prisms Study Group; Lancet 1998, V352, P1498 (29) Prisms Study Group and the University of British Columbia MS/MRI Analysis Group; Neurology 2001, V56, P1628 (30) Runkel, L; Pharm Res 1998, V15, P641 HCAPLUS (31) Salmon, P; J Interferon Cytokine Res 1996, V16, P759 HCAPLUS

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- L66 ANSWER 5 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2002:834389 HCAPLUS
- DN 137:304506
- ED Entered STN: 03 Nov 2002
- TI Pharmacokinetic and pharmacodynamic studies of a human serum albumin-interferon- $\alpha$  fusion protein in cynomolgus monkeys
- AU Osborn, Blaire L.; Olsen, Henrik S.; Nardelli, Bernardetta; Murray, James H.; Zhou, Joe X. H.; Garcia, Andrew; Moody, Gordon; Zaritskaya, Liubov S.; Sung, Cynthia
- CS Human Genome Sciences, Inc., Rockville, MD, USA
- SO Journal of Pharmacology and Experimental Therapeutics (2002), 303(2), 540-548
  CODEN: JPETAB; ISSN: 0022-3565
- PB American Society for Pharmacology and Experimental Therapeutics
- DT Journal
- LA English
- CC 1-7 (Pharmacology)
   Section cross-reference(s): 15
- AB Interferon- $\alpha$  (IFN- $\alpha$ )
  - is indicated for the treatment of certain viral **infections** including hepatitis B and C, and cancers such as melanoma. The short circulating half-life of unmodified **IFN**- $\alpha$  makes frequent dosing (daily or three times weekly) over an extended period (6-12 mo or more) necessary. To improve the pharmacokinetics of **IFN**- $\alpha$  and decrease dosing frequency, **IFN**
  - $-\alpha$  was **fused** to human serum **albumin**
  - producing a new protein, Albuferon. In vitro comparisons of Albuferon and  $\mathbf{IFN}\mathbf{-}\alpha$  showed similar antiviral and
  - antiproliferative activities, although Albuferon was less potent on a molar basis than  $\text{IFN-}\alpha$  . Pharmacokinetic and
  - pharmacodynamic properties of the **fusion** protein were enhanced in monkeys. After a single i.v. injection (30  $\mu$ g/kg) clearance was 0.9 mL/h/kg, and the terminal half-life was 68 h. After 30  $\mu$ g/kg s.c.
  - injection, apparent clearance (clearance divided by bioavailability) was 1.4 mL/h/kg, the terminal half-life was 93 h, and bioavailability was 64%. The rate of clearance of Albuferon was approx. 140-fold slower, and the
  - half-life 18-fold longer, than for  $\mathbf{IFN}-\alpha$  given by the s.c. route in other monkey studies. Sera from Albuferon-treated monkeys demonstrated dose-related antiviral activity for  $\geq 8$  days based on an in vitro bioassay, whereas antiviral activity from  $\mathbf{IFN}-\alpha$ -treated animals was only slightly elevated relative to

vehicle on day 0. Significant increases in 2',5'-oligoadenylate synthetase mRNA relative to  $IFN-\alpha$  - or

- vehicle-treated animals were maintained for ≥10 days after s.c. dosing. The improved pharmacokinetics of Albuferon are accompanied by an improved pharmacodynamic response suggesting that Albuferon may offer the benefits of less frequent dosing and a potentially improved efficacy
- profile compared with  $\mathbf{IFN}$ - $\alpha$ . ST Albuferon interferon antiviral antiproliferative pharmacokinetics pharmacodynamics
- IT Antiviral agents
  Cytotoxic agents
  Human
  Macaca irus

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Pharmacodynamics
     Pharmacokinetics
        (pharmacokinetic and pharmacodynamic studies of a human serum
        albumin-interferon-\alpha fusion
        protein in cynomolgus monkeys)
ΙT
     Albumins, biological studies
     RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (serum, fusion protein with interferon-
        \boldsymbol{\alpha} ; pharmacokinetic and pharmacodynamic studies of a human
        serum albumin-interferon-\alpha
        fusion protein in cynomolgus monkeys)
IT
     Interferons
     RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\alpha , fusion protein with human serum
        albumin; pharmacokinetic and pharmacodynamic studies of a human
        serum albumin-interferon-α
        fusion protein in cynomolgus monkeys)
     69106-44-1, 2',5'-Oligoadenylate synthetase
ΙT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (pharmacokinetic and pharmacodynamic studies of a human serum
        albumin-interferon-α fusion
        protein in cynomolgus monkeys)
                            472960-22-8, Albuferon
IT
     98530-12-2, Intron-A
     RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmacokinetic and pharmacodynamic studies of a human serum
        albumin-interferon-\alpha fusion
        protein in cynomolgus monkeys)
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L66
     2001:781112 HCAPLUS
ΑN
DN
     135:348852
ΕD
     Entered STN: 26 Oct 2001
ΤI
     Albumin fusion proteins with therapeutic proteins for
     improved shelf-life
     Rosen, Craig A.; Haseltine, William A.
ΙN
     Human Genome Sciences, Inc., USA
PA
SO
     PCT Int. Appl., 394 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM C12N015-00
IC
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3, 15
FAN.CNT 7
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
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                                           WO 2001-US11991
                                                            20010412
     WO 2001079480
                       Α1
                            20011025
ΡI
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                      C2
                            20030109
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                       A1
                            20030122
                                           EP 2001-937179
                                                           20010412
     EP 1276856
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     US 2003125247
                      Α1
                            20030703
                                           US 2001-833041
                                                            20010412
                            20030911
                                           US 2001-833117
     US 2003171267
                       A1
                                                            20010412
     JP 2003530852
                       T2
                            20031021
                                           JP 2001-577463
                                                            20010412
                                           US 2001-832501
     US 2003199043
                       Α1
                            20031023
                                                            20010412
                                           US 2001-833118
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                       Α1
                            20031127
                                                            20010412
                                           US 2001-833245
     US 2004010134
                       Α1
                            20040115
                                                            20010412
PRAI US 2000-229358P
                       Ρ
                            20000412
     US 2000-199384P
                       Ρ
                            20000425
     US 2000-256931P
                       Ρ
                            20001221
                       W . 20010412
     WO 2001-US11991
     The present invention encompasses fusion proteins of
AΒ
     albumin with various therapeutic proteins. Therapeutic proteins
     may be stabilized to extend the shelf-life, and/or to
     retain the therapeutic protein's activity for extended periods of time in
     solution, in vitro and/or in vivo, by genetically or chemical fusing
     or conjugating the therapeutic protein to albumin or a fragment
     or variant of albumin. Use of albumin fusion
     proteins may also reduce the need to formulate the protein solns. with
     large excesses of carrier proteins to prevent loss of therapeutic proteins
     due to factors such as binding to the container. Nucleic acid mols.
     encoding the albumin fusion proteins of the invention
     are also encompassed by the invention, as are vectors containing these nucleic
     acids, host cells transformed with these nucleic acids vectors, and
     methods of making the albumin fusion proteins of the
     invention and using these nucleic acids, vectors, and/or host cells.
     Thus, plasmid vectors are constructed in which DNA encoding the desired
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therapeutic protein may be inserted for expression of the albumin fusion proteins in yeast (pPPC0005) and mammalian cells (pC4:HSA). Yeast-derived signal sequences from Saccharomyces cerevisiae invertase SUC2 gene, or the stanniocalcin or native human serum albumin signal peptides, are used for secretion in yeast or mammalian systems, resp. Thus, the fusion product of human growth hormone with residues 1-387 of human serum albumin retains essentially intact biol. activity after 5 wk of incubation in tissue culture media at 37°, whereas recombinant human growth hormone used as control lost its biol. activity in the first week. Although the potency of the albumin fusion proteins is slightly lower than the unfused counterparts in rapid bioassays, their biol. stability results in much higher biol. activity in the longer term in vitro assay or in vivo Addnl., the present invention encompasses pharmaceutical compns. comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disorders or conditions using albumin fusion proteins of the invention. albumin fusion therapeutic protein shelflife Receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (4-1BB; albumin fusion proteins with therapeutic proteins for improved shelf-life) Cytokines RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (BAFF; albumin fusion proteins with therapeutic proteins for improved shelf-life) Cytokine receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (DR4 (death receptor 4); albumin fusion proteins with therapeutic proteins for improved shelf-life) Cytokine receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (DR5 (death receptor 5); albumin fusion proteins with therapeutic proteins for improved shelf-life) Cytokines RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (MPIF-1 (myeloid progenitor inhibitory factor 1); albumin fusion proteins with therapeutic proteins for improved shelf-life) Steroid receptors Thyroid hormone receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (TR (thyroid/steroid hormone receptor), 11; albumin fusion proteins with therapeutic proteins for improved shelf-life) Steroid receptors Thyroid hormone receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (TR (thyroid/steroid hormone receptor), 12; albumin fusion proteins with therapeutic proteins for improved shelf-life) Steroid receptors Thyroid hormone receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(TR (thyroid/steroid hormone receptor), 13; albumin

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fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Steroid receptors
    Thyroid hormone receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TR (thyroid/steroid hormone receptor), 14; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Steroid receptors
    Thyroid hormone receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TR (thyroid/steroid hormone receptor), 16; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Steroid receptors
    Thyroid hormone receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TR (thyroid/steroid hormone receptor), 8; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Steroid receptors
ΙT
    Thyroid hormone receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TR2 (thyroid/steroid hormone receptor 2); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Steroid receptors
    Thyroid hormone receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TR3 (thyroid/steroid hormone receptor 3); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TRAIL (tumor necrosis factor-related apoptosis-inducing ligand);
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
    Cytokine receptors
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TRAIL, 4; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Cytokine receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TRAIL, 6; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Cytokine receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TRAIL-R3; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Drug delivery systems
    Gene therapy
    Molecular cloning
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
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مه واليتان

والمتناث

و بالتيزاد

ور ونيته

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Cell adhesion molecules
TT
     Cytokines
     Enzymes, biological studies
     Fas antigen
     Fas ligand
       Fusion proteins (chimeric proteins)
     Growth factors, animal
       Interferons
     Synthetic gene
     Tumor necrosis factor receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     Proteins, specific or class
TΤ
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (apoptosis-regulating, AIM-2; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Cytokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (endokine; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Signal peptides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (for improved secretion in yeast or mammalian cells; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
     Interferons
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (keratinocyte-derived; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TT
     Animal cell
        (mammalian, recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT.
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
     Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Plasmid vectors
        (pScCHSa, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
TΤ
     Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
ΙT
     Saccharomyces cerevisiae
     Yeast
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
     Albumins, biological studies
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RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic

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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (serum; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
         (signal sequence, for improved secretion in yeast or mammalian cells;
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
ΙT
     Antibodies
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (single chain; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (therapeutic; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
      Interferons
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (\alpha ; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TT
     Chemokine receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (β chemokine receptor CCR5; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Tumor necrosis factors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
             albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
     Tumor necrosis factors
TT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (\delta; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
      189460-40-0P, Connective tissue growth factor
TΤ
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (2 and 4; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
      9001-84-7P, Phospholipase A2
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (T-cell lymphoma lipoprotein-associated; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
                                        9002-68-0P, FSH
                                                           9002-72-6P, Growth
IΤ
      9002-67-9P, Luteinizing hormone
                9004-10-8P, Insulin, biological studies
                                                          11096-26-7P,
     hormone
                       67763-96-6P, Insulin-like growth factor 1
                                                                    83869-56-1P,
     Erythropoietin
              124861-55-8P, Proteinase inhibitor, TIMP-2
     GM-CSF
      127464-60-2P, Vascular endothelial growth factor 140208-24-8P,
                                     143011-72-7P, G-CSF
      Proteinase inhibitor, TIMP-1
      145809-21-8P, Proteinase inhibitor, TIMP-3
                                                   148348-15-6P,
                                   171758-70-6P, Keratinocyte growth factor 2
      Fibroblast growth factor 7
     186207-03-4P, Proteinase inhibitor, TIMP-4
                                                   205944-50-9P,
                                                                244019-42-9P,
                        207621-35-0P, Osteoprotegerin ligand
      Osteoprotegerin
     Vascular endothelial growth factor 2
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
    127361-02-8DP, Albumin (human blood serum clone HSA-II/HSA-I-A
IT
    protein moiety reduced), full-length or subfragment fusion
    products
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    155945-98-5, PN: US5962255 SEQID: 59 unclaimed DNA
                                                          156163-00-7
                                             167728-72-5 167728-73-6
    167728-69-0
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                                167728-71-4
                  167731-74-0, PN: US5962255 SEQID: 56 unclaimed DNA
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                    167731-78-4, PN: US5962255 SEQID: 61 unclaimed DNA
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    US5962255 SEQID: 551 unclaimed DNA
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                                                        167732-13-0
    167732-14-1, PN: US5962255 SEQID: 554 unclaimed DNA
                                                          167732-15-2, PN:
                                        167732-16-3 167732-17-4
    US5962255 SEQID: 555 unclaimed DNA
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    167732-20-9, PN: US5962255 SEQID: 572 unclaimed DNA
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    167732-22-1, PN: US5962255 SEQID: 574 unclaimed DNA
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                                   217893-78-2, GenBank A63615
    217893-77-1, GenBank A63614
                                                                 217893-79-3,
                     217893-80-6, GenBank A63617
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    GenBank A63616
                                  217893-83-9, GenBank A63620
                                                                 217893-84-0,
    217893-82-8, GenBank A63619
    GenBank A63621
                     217893-85-1, GenBank A63622
                                                   217893-86-2, GenBank A63624
    217893-89-5, GenBank A63627 217893-90-8, GenBank A63628
                                                                 217893-91-9,
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                  367319-55-9
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    367319-59-3
                   367319-60-6
                                 367319-61-7
                                367319-66-2
    367319-64-0
                   367319-65-1
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
       proteins with therapeutic proteins for improved shelf-
        life)
                                               352583-76-7, Protein (human
ΙT
     173586-11-3
                  221879-28-3
                                 222614-92-8
     clone 785CIP2B 67)
                         370649-84-6
                                      370649-85-7
     RL: PRP (Properties)
        (unclaimed protein sequence; albumin fusion
       proteins with therapeutic proteins for improved shelf-
        life)
                                244008-03-5, PN: WO9947540 SEQID: 3 unclaimed
                   131748-18-0
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     122024-47-9
          244008-06-8, PN: WO9947540 SEQID: 4 unclaimed DNA 244008-07-9, PN:
                                        244008-08-0, PN: WO9947540 SEQID: 6
    WO9947540 SEQID: 5 unclaimed DNA
                    244008-09-1, PN: WO9947540 SEQID: 7 unclaimed DNA
    unclaimed DNA
     244008-12-6, 8: PN: WO0183510 SEQID: 8 unclaimed DNA
                                                           244008-13-7, PN:
    WO9947540 SEQID: 9 unclaimed DNA 244008-14-8, PN: WO9947540 SEQID: 10
                    367273-46-9
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                                                             370598-71-3
    unclaimed DNA
     370649-86-8
    RL: PRP (Properties)
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
             THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Delta Biotechnology Limited; EP 0322094 Al 1989 HCAPLUS
(2) Delta Biotechnology Limited; WO 9523857 Al 1995 HCAPLUS
    ANSWER 7 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
ΑN
     2001:781079 HCAPLUS
   . 135:348851
DN
    Entered STN: 26 Oct 2001
ED
    Albumin fusion proteins with therapeutic proteins for
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improved shelf-life
IN
    Rosen, Craig A.; Haseltine, William A.
     Human Genome Sciences, Inc, USA
PΑ
SO
     PCT Int. Appl., 606 pp.
     CODEN: PIXXD2
DT
     Patent
LA
    English
IC
     ICM C12N
CC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3, 15
FAN.CNT 7
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
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     WO 2001079444
                      A2
                            20011025
                                           WO 2001-US12013 20010412
     WO 2001079444
                     A3
                            20020523
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
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             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
    AU 2001074809
                     A5
                            20011020
                                          AU 2001-74809
                                                           20010412
     EP 1278544
                       Α2
                            20030129
                                           EP 2001-941457 20010412
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     US 2003125247
                      Α1
                            20030703
                                           US 2001-833041
                                                            20010412
     US 2003171267
                            20030911
                                           US 2001-833117
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                       Α1
     JP 2003530847
                      Т2
                            20031021
                                           JP 2001-577428
                                                            20010412
    US 2003199043
                     Α1
                            20031023
                                           US 2001-832501
                                                            20010412
     US 2003219875
                     Α1
                            20031127
                                           US 2001-833118
                                                            20010412
     US 2004010134
                      A1
                            20040115
                                           US 2001-833245
                                                            20010412
PRAI US 2000-229358P
                      Ρ
                            20000412
    US 2000-199384P
                      Ρ
                            20000425
     US 2000-256931P
                      Ρ
                            20001221
    WO 2001-US12013
                     W
                            20010412
     The present invention encompasses fusion proteins of
AΒ
    albumin with various therapeutic proteins. Therapeutic proteins
    may be stabilized to extend the shelf-life, and/or to
     retain the therapeutic protein's activity for extended periods of time in
     solution, in vitro and/or in vivo, by genetically or chemical fusing
    or conjugating the therapeutic protein to albumin or a fragment
     or variant of albumin. Use of albumin fusion
    proteins may also reduce the need to formulate the protein solns. with
     large excesses of carrier proteins to prevent loss of therapeutic proteins
     due to factors such as binding to the container. Nucleic acid mols.
     encoding the albumin fusion proteins of the invention
     are also encompassed by the invention, as are vectors containing these nucleic
     acids, host cells transformed with these nucleic acids vectors, and
    methods of making the albumin fusion proteins of the
     invention and using these nucleic acids, vectors, and/or host cells.
     Thus, plasmid vectors are constructed in which DNA encoding the desired
     therapeutic protein may be inserted for expression of the albumin
     fusion proteins in yeast (pPPC0005) and mammalian cells (pC4:HSA).
     Yeast-derived signal sequences from Saccharomyces cerevisiae invertase
     SUC2 gene, or the stanniocalcin or native human serum albumin
     signal peptides, are used for secretion in yeast or mammalian systems,
     resp. Thus, the fusion product of human growth hormone with
     residues 1-387 of human serum albumin retains essentially intact
    biol. activity after 5 wk of incubation in tissue culture media at
     37°, whereas recombinant human growth hormone used as
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control lost its biol. activity in the first week. Although the potency
of the albumin fusion proteins is slightly lower than
the unfused counterparts in rapid bioassays, their biol. stability results
in much higher biol. activity in the longer term in vitro assay or in vivo
       Addnl., the present invention encompasses pharmaceutical compns.
comprising albumin fusion proteins and methods of
treating, preventing, or ameliorating diseases, disorders or conditions
using albumin fusion proteins of the invention.
albumin fusion therapeutic protein shelflife
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (1-309; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (11; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (12; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (15; albumin fusion proteins with therapeutic
  proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (17; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (18; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Interleukins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (19; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (1; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Interleukins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (21; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Bone morphogenetic proteins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (2; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
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(331D5; albumin fusion proteins with therapeutic

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proteins for improved shelf-life)
    Bone morphogenetic proteins
TΤ
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (3; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (4-1BB; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    Bone morphogenetic proteins
IT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (4; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Bone morphogenetic proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (5; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (61164; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Bone morphogenetic proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (6; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Bone morphogenetic proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (7; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Bone morphogenetic proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (9; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Platelet-derived growth factors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (AA; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ACRP-30; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ADEC (adenoid expressed chemokine); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Interleukins
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (AGF; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
     Proteins, specific or class
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RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (APM-1; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (Act-2; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Platelet-derived growth factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (BB; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (BCMA; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Platelet-derived growth factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (Bv-sis; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, 2; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, 3; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, DGWCC; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, DVic-1; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, ELC; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, HCC-1; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, IBICK; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (C-C, ILINCK; albumin fusion proteins with
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therapeutic proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C-C, SLC (secondary lymphoid chemokine); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C-C, STCP-1; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C-X-C, 3; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C-X-C; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C10; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
ΙT
    Troponins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (C; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CCC3; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); .PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CCF18; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CCR2; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
IT
    CD antigens
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CD27; albumin fusion proteins with therapeutic
       proteins for improved shelf-life)
IT
    Glycoproteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CD40-L (antigen CD40 ligand); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CTAP-III (connective tissue activating protein III); albumin
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fusion proteins with therapeutic proteins for improved

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shelf-life)
ΙT
    Antigens
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CTLA-8; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Chemokine receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (CXCR3; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Cerebus; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Chr19Kine; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Platelet-derived growth factors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (D; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Cytokine receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (DR3 (death receptor 3); albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
     Proteins, specific or class
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (EDAR; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Interleukins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (EDIRF I protein; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
    Chemokines
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (EEC (eosinophil expressed chemokine); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ENA-78 (epithelial neutrophil activating protein-78); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Hemopoietins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (FLT3 ligand; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (HCC-1; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
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ΙT
    Troponins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (I; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (L105-7; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (LVEC-1 (liver expressed chemokine 1); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (LVEC-2 (liver expressed chemokine 2); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
    Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Lyn-1; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (M110; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (M11A; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (MACK (mammary associated chemokine); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (MCP-3\alpha and MCP-3\beta;
                              albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (MCP-4; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
TΤ
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (MCPP (monocyte chemotactic proprotein); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(MDC (macrophage-derived chemokine); albumin fusion
   proteins with therapeutic proteins for improved shelf-
Monokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (MIG (monokine induced by \gamma- interferon);
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (MIG-\beta; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Interleukins
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (MIRAP; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (MP52; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (NOGO-66; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (NOGO-A; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (NOGO-B; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (NOGO-C; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Antigens
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (OX-40; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (PF4; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (PGBC (pituitary expressed chemokine); albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Chemokine receptors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(RANTES; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (SISD; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (SLC (secondary lymphoid tissue chemokine); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Troponins
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (T; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TAC1; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Cytokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TARC (thymus and activation regulated cytokine); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TMEC (T cell mixed lymphocyte reaction expressed chemokine);
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Tarc; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Tim-1; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Troy; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
TΤ
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ZCHEMO-8; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ZSIG-35; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Drug delivery systems
     Gene therapy
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Molecular cloning

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(albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
ΙΤ
    CD30 (antigen)
    CD40 (antigen)
    Cell adhesion molecules
    Cytokines
    Enzymes, biological studies
    Eotaxin
    Erythropoietin receptors
    Fas ligand
       Fusion proteins (chimeric proteins)
    Granulocyte-macrophage colony-stimulating factor receptors
    Growth factors, animal
       Interferons
    Interleukin 1
    Interleukin 1 receptor antagonist
    Interleukin 11
    Interleukin 13
    Interleukin 14
    Interleukin 15
    Interleukin 17
    Interleukin 18
    Interleukin 1\alpha
    Interleukin 1B
    Interleukin 3
    Interleukin 4
    Interleukin 4 receptors
    Interleukin 5 receptors
    Interleukin 6
    Interleukin 6 receptors
    Interleukin 8
    Interleukin 8 receptors
    Interleukin 9
    Lymphotoxin
    Monocyte chemoattractant protein-1
    Neutrophil-activating peptide-2
    Platelet-derived growth factors
    RANTES (chemokine)
    Stem cell factor
    Synthetic gene
    Tumor necrosis factor receptors
    Tumor necrosis factors
    Vascular endothelial growth factor receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
ΙT
    Interleukin 10
     Interleukin 12
    Interleukin 2
    Interleukin 5
    Interleukin 7
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
IT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (b57; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
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(chemokine-like protein PF4-414; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Growth factors, animal
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (chondromodulins, -like protein; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Proteins, specific or class
ΙT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (collapsins, antibodies for; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
TT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (exodus; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Signal peptides
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (for improved secretion in yeast or mammalian cells; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
     Chemokines
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (fractalkines; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Agglutinins and Lectins
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (galectin-4; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene Patched-2; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TΨ
     Vascular endothelial growth factor receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene flt 1; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     Vascular endothelial growth factor receptors
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene flt 4; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     Proteins, specific or class
IΤ
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gene patched; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (glycodelin-A; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Chemokines
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RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic

use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(granulocyte chemotactic protein-2; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gro-\alpha; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gro-\beta; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (gro-γ; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     Proteins, specific or class
ΙT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (growth-related oncogene-\alpha; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Proteins, specific or class
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (growth-related oncogene-\beta; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Proteins, specific or class
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (growth-related oncogene-γ; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT'
     Cytokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interferon-inducible IP-10; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Interleukin receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interleukin 10 receptors; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
     Interleukin receptors
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interleukin 11; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Interleukin receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interleukin 12; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Interleukin receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interleukin 13; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Interleukin receptors
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RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin 15; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Interleukin receptors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin 17; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Interleukin receptors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin 9; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin C; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin-1 accessory; albumin fusion proteins
   with therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (interleukin-2 receptor associated p43; albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Lymphokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (lymphotactins; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (macrophage inflammatory protein 3\alpha; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (macrophage inflammatory protein 3\beta; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (macrophage inflammatory protein 3\gamma; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Animal cell
   (mammalian, recombinant expression host; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Antitumor agents
   (melanoma; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Chemokines
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(monocyte chemoattractant protein 3; albumin fusion
        proteins with therapeutic proteins for improved shelf-
ΙT
     Chemokine receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (monocyte chemoattractant protein-1; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Chemokines
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (monocyte chemoattractant protein-2; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Chemokine receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (monocyte chemoattractant protein-4; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Proteins, specific or class
ΙT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (neurotactin; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Growth factors, animal
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (osteogenic protein 2; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Tumor necrosis factor receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (p75; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     Plasmid vectors
        (pScCHSa, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
TΤ
     Placental hormones
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (placenta-derived mitogenic factors; albumin fusion
        proteins with therapeutic proteins for improved shelf-
ΙT
     Saccharomyces cerevisiae ·
     Yeast
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
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shelf-life)

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Albumins, biological studies
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (serum; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
TT
    Genetic element
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (signal sequence, for improved secretion in yeast or mammalian cells;
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
    Antibodies
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (single chain; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TТ
    Chemokines
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (stem cell inhibitory factor; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
    Growth factors, animal
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (stroma-derived growth factor 1\alpha and 1\beta;
                                                   albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Proteins, specific or class
TT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (therapeutic; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Interleukin 1 receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (type 3; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Interleukin 1 receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (type II; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Interferons
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (\alpha ; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
TΤ
    Chemokine receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (β chemokine receptor CCR5; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Chemokine receptors
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (β chemokine receptor CCR7; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Transforming growth factors
ΤТ
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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(\beta 1-; albumin fusion proteins with therapeutic
         proteins for improved shelf-life)
IT
     Transforming growth factors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (\beta 2-; albumin fusion proteins with therapeutic
         proteins for improved shelf-life)
·IT
     Chemokines
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (\beta 9; albumin fusion proteins with therapeutic
         proteins for improved shelf-life)
 ΙT
     Thrombomodulin
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (\beta; albumin fusion proteins with therapeutic
         proteins for improved shelf-life)
ΙT
     78990-62-2P, Calpain
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (10a and 10b and 10c; albumin fusion proteins with
         therapeutic proteins for improved shelf-life)
                                               9002-62-4P, Prolactin, biological
 ΙT
     50-56-6P, Oxytocin, biological studies
                9002-67-9P, Luteinizing hormone 9002-68-0P, FSH
                                                                    9002-72-6P,
                       9004-10-8P, Insulin, biological studies
                                                                 9014-42-0P,
     Growth hormone
                       11000-17-2P, Vasopressin
                                                 11096-26-7P, Erythropoietin
     Thrombopoietin
     33507-63-0P, Substance P
                                 67763-96-6P, Insulin-like growth factor 1
                           106096-92-8P, Acidic fibroblast growth factor
     83869-56-1P, GM-CSF
     106096-93-9P, Basic fibroblast growth factor
                                                     122191-40-6P, ICE
                  123584-45-2P, Fibroblast growth factor 4
                                                              129653-64-1P,
     proteinase
                                   130939-41-2P, Fibroblast growth factor 6
     Fibroblast growth factor 5
     130939-66-1P, Neurotrophin 3
                                   140208-23-7P, Plasminogen activator
                   141760-45-4P, Furin
                                          142243-03-6P, Plasminogen activator
     inhibitor-1
                                          143375-33-1P, Neurotrophin 4
                    143011-72-7P, G-CSF
     inhibitor-2
     148348-14-5P, Fibroblast growth factor 3
                                                 151185-16-9P, Fibroblast growth
                157857-21-1P, Maspin
                                       164003-41-2P, Fibroblast growth factor 8
     185915-22-4P, Fibroblast growth factor 13
                                                  187888-07-9P, Endostatin
     193363-12-1P, Vascular endothelial growth factor D
                                                          203874-76-4P,
                                   204719-95-9P, Fibroblast growth factor 16
     Fibroblast growth factor 12
                                   219563-02-7P, Vascular endothelial growth
     214210-47-6P, Neuropilin 1
                                              271597-10-5P,
                227018-38-4P, Neuropilin 2
     Growth/differentiation factor 1
                                        322637-18-3P, Fibroblast growth factor
                                    332350-92-2P, Bone morphogenetic protein
           331718-56-0P, Resistin
     receptor kinase 3
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (albumin fusion proteins with therapeutic proteins
         for improved shelf-life)
 IT
     144114-21-6, Retropepsin
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
         (inhibitors; albumin fusion proteins with
         therapeutic proteins for improved shelf-life)
 IT
     127464-60-2P, Vascular endothelial growth factor
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (isoforms; albumin fusion proteins with therapeutic
         proteins for improved shelf-life)
     127361-02-8DP, Albumin (human blood serum clone HSA-II/HSA-I-A
 IT
     protein moiety reduced), full-length or subfragment fusion
     products
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
```

(nucleotide sequence; albumin fusion proteins with

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therapeutic proteins for improved shelf-life)
     155945-98-5, PN: US5962255 SEQID: 59 unclaimed DNA
                                                          156163-00-7
IT
                   167728-70-3
                                              167728-72-5
     167728-69-0
                                 167728-71-4
                                                             167728-73-6
     167731-70-6
                   167731-74-0, PN: US5962255 SEQID: 56 unclaimed DNA
     167731-75-1, PN: US5962255 SEQID: 57 unclaimed DNA
                                                          167731-76-2, PN:
     US5962255 SEQID: 58 unclaimed DNA
                                       167731-77-3, PN: US5962255 SEQID: 60
                     167731-78-4, PN: US5962255 SEQID: 61 unclaimed DNA
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                                 167731-81-9
                                              167732-10-7
     167731-79-5
                   167731-80-8
                                                             167732-11-8, PN:
                                         167732-12-9
     US5962255 SEQID: 551 unclaimed DNA
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     US5962255 SEQID: 555 unclaimed DNA 167732-16-3
                                                        167732-17-4
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     167732-18-5
     167732-20-9, PN: US5962255 SEQID: 572 unclaimed DNA
                                                          167732-21-0
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                                                           195164-37-5
     217893-77-1, GenBank A63614
                                   217893-78-2, GenBank A63615
                                                                 217893-79-3,
     GenBank A63616
                      217893-80-6, GenBank A63617
                                                    217893-81-7, GenBank A63618
     217893-82-8, GenBank A63619
                                   217893-83-9, GenBank A63620
                                                                 217893-84-0,
                                                    217893-86-2, GenBank A63624
     GenBank A63621
                      217893-85-1, GenBank A63622
     217893-89-5, GenBank A63627
                                   217893-90-8, GenBank A63628
                                                                 217893-91-9,
                      217893-92-0, GenBank A63630 244008-03-5, PN: WO9947540
     GenBank A63629
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                              367319-52-6
                                           367319-53-7
                                                          367319-54-8
     367319-55-9
                   367319-56-0
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                                               367319-58-2
                                                             367319-59-3
                                 367319-62-8
                                               367319-63-9
                                                             367319-64-0
     367319-60-6
                   367319-61-7
                                 370965-07-4
                                               370965-08-5
     367319-65-1
                   367319-66-2
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
                   131748-18-0
                                 244008-06-8, PN: WO9947540 SEQID: 4 unclaimed
     122024-47-9
           244008-07-9, PN: WO9947540 SEQID: 5 unclaimed DNA
                                                               244008-08-0, PN:
     WO9947540 SEQID: 6 unclaimed DNA 244008-09-1, PN: WO9947540 SEQID: 7
                    244008-12-6, 8: PN: WO0183510 SEQID: 8 unclaimed DNA
     unclaimed DNA
     244008-13-7, PN: WO9947540 SEQID: 9 unclaimed DNA
                                                       367273-46-9
                                371149-71-2
     367273-47-0
                  367273-48-1
     RL: PRP (Properties)
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     102510-92-9P, Inhibin A
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (\alpha- and \beta-subunits;
                             albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
     9061-61-4P, Nerve growth factor
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (β; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    ANSWER 8 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
     2001:781078 HCAPLUS
AN
DΝ
     135:348850
     Entered STN: 26 Oct 2001
ED
     Albumin fusion proteins with therapeutic proteins for
TI
     improved shelf-life
IN
     Rosen, Craig A.; Haseltine, William A.
PΑ
     Human Genome Sciences, Inc., USA
SO
     PCT Int. Appl., 374 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM C12N
CC
     63-3 (Pharmaceuticals)
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Section cross-reference(s): 3, 15

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     PATENT NO.
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                                          WO 2001-US11924 20010412 ·
     WO 2001079443
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                     А3
                           20020221
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
                    CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
             BJ, CF,
                                          AU 2001-59063
                                                           20010412
     AU 2001059063
                      Α5
                            20011030
     EP 1274719
                      A2
                           20030115
                                          EP 2001-932546
                                                           20010412
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                           US 2001-833041
                                                            20010412
     US 2003125247
                      Α1
                            20030703
                                          US 2001-833117
     US 2003171267
                      Α1
                           20030911
                                                            20010412
     JP 2003530846
                      Т2
                           20031021
                                          JP 2001-577427
                                                           20010412
     US 2003199043
                      Α1
                           20031023
                                          US 2001-832501
                                                            20010412
     US 2003219875
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                          20031127
                                          US 2001-833118
                                                            20010412
     US 2004010134
                      A1
                           20040115
                                           US 2001-833245
                                                            20010412
PRAI US 2000-229358P P
                           20000412
     US 2000-199384P P
                            20000425
     US 2000-256931P
                     Ρ
                            20001221
     WO 2001-US11924
                     W
                           20010412
AB
     The present invention encompasses fusion proteins of
     albumin with various therapeutic proteins. Therapeutic proteins
     may be stabilized to extend the shelf-life, and/or to
     retain the therapeutic protein's activity for extended periods of time in
     solution, in vitro and/or in vivo, by genetically or chemical fusing
     or conjugating the therapeutic protein to albumin or a fragment
     or variant of albumin. Use of albumin fusion
     proteins may also reduce the need to formulate the protein solns. with
     large excesses of carrier proteins to prevent loss of therapeutic proteins
     due to factors such as binding to the container. Nucleic acid mols.
     encoding the albumin fusion proteins of the invention
     are also encompassed by the invention, as are vectors containing these nucleic
     acids, host cells transformed with these nucleic acids vectors, and
     methods of making the albumin fusion proteins of the
     invention and using these nucleic acids, vectors, and/or host cells.
     Thus, plasmid vectors are constructed in which DNA encoding the desired
     therapeutic protein may be inserted for expression of the albumin
     fusion proteins in yeast (pPPC0005) and mammalian cells (pC4:HSA).
     Yeast-derived signal sequences from Saccharomyces cerevisiae invertase
     SUC2 gene, or the stanniocalcin or native human serum albumin
     signal peptides, are used for secretion in yeast or mammalian systems,
     resp. Thus, the fusion product of human growth hormone with
     residues 1-387 of human serum albumin retains essentially intact
     biol. activity after 5 wk of incubation in tissue culture media at
     37°, whereas recombinant human growth hormone used as
     control lost its biol. activity in the first week. Although the potency
     of the albumin fusion proteins is slightly lower than
     the unfused counterparts in rapid bioassays, their biol. stability results
     in much higher biol. activity in the longer term in vitro assay or in vivo
             Addnl., the present invention encompasses pharmaceutical compns.
     comprising albumin fusion proteins and methods of
     treating, preventing, or ameliorating diseases, disorders or conditions
     using albumin fusion proteins of the invention.
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albumin fusion therapeutic protein shelflife

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IT
    Bone morphogenetic proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (2; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    Bone morphogenetic proteins
IT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (7; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Transport proteins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ABC1 (ATP-binding cassette-containing 1); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Proteins, specific or class
IT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (ADMP (anti-dorsalizing morphogenetic protein-1); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Agouti signal; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BPI (bactericidal/permeability-increasing), 21; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Transcription factors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BRCA1; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
    Transcription factors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (BRCA2; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    Proteins, specific or class
IT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (Del-1 (developmentally regulated endothelial locus-1); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Proteins, specific or class
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (EMAP II (endothelial monocyte activating polypeptide II);
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
ΙT
    Troponins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (I; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
     Toxins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)

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Myelin basic protein

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(ML-I (mistletoe lectin I); albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (MTP (microsomal transfer protein); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙΤ
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (NIF (neutrophil inhibitory factor); albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΤТ
    Receptors
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (T1/ST2; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙΤ
    Glycoproteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TNF-BP (tumor necrosis factor-binding protein); albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (TRAIL (tumor necrosis factor-related apoptosis-inducing ligand);
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
IΤ
     Drug delivery systems
     Gene therapy.
    Molecular cloning
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
IT
    Arrestins
    CD4 (antigen)
    CTLA-4 (antigen)
    Calreticulin
    Cell adhesion molecules
    Ciliary neurotrophic factor
    Cytokines
    Decorins
    Enzymes, biological studies
       Fusion proteins (chimeric proteins)
    Gelsolin
    Growth factors, animal
     Heat-shock proteins
       Interferons
     Interleukin 1
     Interleukin 1 receptor antagonist
     Interleukin 10
     Interleukin 11
     Interleukin 12
     Interleukin 18
     Interleukin 4
     Interleukin 4 receptors
     Interleukin 8
    LFA-3 (antigen)
    Lactoferrins
     Leukemia inhibitory factor
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Platelet-derived growth factors
Pleiotrophins
Stem cell factor
Synthetic gene
Tumor necrosis factor receptors
Tumor necrosis factor receptors
Tumor necrosis factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (albumin fusion proteins with therapeutic proteins
   for improved shelf-life)
Neurotrophic factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (brain-derived; albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (chemokine-binding; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (corticotropin-releasing factor-binding; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (diphtheria, fusion protein with interleukin 2;
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (exotoxins, Pseudomonas, fusion protein with acidic
   fibroblast growth factor; albumin fusion proteins
   with therapeutic proteins for improved shelf-life)
Signal peptides
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
   (for improved secretion in yeast or mammalian cells; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Interleukin 3
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (fusion protein with G-CSF; albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Interleukin 6
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (fusion proteins with diphtheria toxin or Pseudomonas
   exotoxin; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (gene patched; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
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Neurotrophic factors
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (glial-derived; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Interferons
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interferon \omega; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Proteins, specific or class
ΙT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (interferon-induced, 10; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Animal cell
        (mammalian, recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Proteins, specific or class
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (noggins; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; {\bf albumin}
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
     Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Plasmid vectors
        (pScCHSa, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
     Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
ΙT
    Hemopoietins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (progenipoietin; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Hemopoietins
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (promegapoietin; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Saccharomyces cerevisiae
     Yeast
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
     Antigens
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (retinal S-; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
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Albumins, biological studies

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RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (serum; albumin fusion proteins with therapeutic
  proteins for improved shelf-life)
Genetic element
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
   (signal sequence, for improved secretion in yeast or mammalian cells;
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
Antibodies
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (single chain; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Hedgehog protein
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (sonic; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (therapeutic; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (tie-2; albumin fusion proteins with therapeutic
  proteins for improved shelf-life)
Complement receptors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (type 1; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Collagens, biological studies
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (type II; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Interferons
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Interferons
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (\alpha ; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Transforming growth factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (\beta 1-; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Transforming growth factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (\beta 2-; albumin fusion proteins with therapeutic
   proteins for improved shelf-life)
Transforming growth factors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
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(β3-; albumin fusion proteins with therapeutic proteins for improved **shelf-life**) IT Interferons RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  $(\gamma)$ ; albumin fusion proteins with therapeutic proteins for improved **shelf-life**) 139691-92-2P, Serine proteinase inhibitor IT RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (1; albumin fusion proteins with therapeutic proteins for improved shelf-life) 9001-91-6DP, Lys-plasminogen, de-(1-76) derivs. IT RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (Lys-plasminogen; albumin fusion proteins with therapeutic proteins for improved shelf-life) 9002-12-4P, 9001-42-7P,  $\alpha$ -Glucosidase 9002-01-1P, Streptokinase ΙT 9002-67-9P, 9002-61-3P, Chorionic gonadotropin Urate oxidase 9002-72-6P, 9002-68-0P, FSH 9002-69-1P, Relaxin Luteinizing hormone Growth hormone 9003-98-9P, DNase 9004-10-8P, Insulin, biological 9007-92-5P, Glucagon, biological studies 9014-42-0P, 9025-35-8P, 9015-68-3P, Asparaginase Thrombopoietin 9026-93-1P, Adenosine deaminase 9035-55-6P, α-Galactosidase 9039-53-6P, Urokinase 9040-61-3P, Staphylokinase 9054-89-1DP, Superoxide dismutase, fusion protein with botulin 9061-61-4P, Nerve growth factor 9073-56-7P,  $\alpha$ -L-Iduronidase 11096-26-7P, Erythropoietin 9088-41-9P, Kunitz proteinase inhibitor 42616-25-1P, Methioninase 37228-64-1P, β-Glucocerebrosidase 55354-43-3P, Arylsulfatase B 62229-50-9P, Epidermal growth factor 76901-00-3P, Platelet 67763-96-6P, Insulin-like growth factor 1 82707-54-8P, Neprilysin 83652-28-2P, activating factor acetylhydrolase Calcitonin gene-related peptide 83869-56-1P, GM-CSF 86090-08-6P, 104625-48-1P, Activin A Angiostatin 99149-95-8P, Saruplase 106096-92-8DP, Acidic 105844-41-5P, Plasminogen activator inhibitor fibroblast growth factor, fusion protein with Pseudomonas 106096-92-8P 106096-93-9P, Fibroblast growth factor 2 exotoxin 107231-12-9DP, Botulin, fusion protein with superoxide dismutase 116036-70-5P, Fibrolase 130939-66-1P, Neurotrophin 3 139639-23-9P, Tissue-type plasminogen activator 143011-72-7P, G-CSF 145137-38-8P, 157857-21-1P, Maspin Desmoteplase 153858-68-5P, Contortrostatin 169494-85-3P, Leptin 186270-49-5P, 163658-39-7P, Prosaptide 194368-66-6P, Angiopoietin 2 194554-71-7P, Tissue Angiopoietin 1 factor pathway inhibitor 195009-21-3P, Glial growth factor 2 197980-93-1P, Pigment epithelium-derived factor 196488-72-9P, Ranpirnase 205944-50-9P, Osteoprotegerin 244019-30-5P, Vascular endothelial growth 362605-29-6P, Keratinocyte growth 320336-96-7P, Kistrin factor 1 factor 1 RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT 9000-95-7P, Apyrase RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (ecto-; albumin fusion proteins with therapeutic proteins for improved shelf-life) 9002-79-3P, MSH IT RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (fusion products with diphtheria toxin; albumin fusion proteins with therapeutic proteins for improved shelf-life)

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مربوانيتني

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ΙT
    127361-02-8DP, Albumin (human blood serum clone HSA-II/HSA-I-A
    protein moiety reduced), full-length or subfragment fusion
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
                               217893-77-1, GenBank A63614
                                                              217893-78-2,
ΙT
                  156163-00-7
     GenBank A63615
                     217893-79-3, GenBank A63616
                                                   217893-80-6, GenBank A63617
                                 217893-82-8, GenBank A63619
     217893-81-7, GenBank A63618
                                                                217893-83-9,
     GenBank A63620
                     217893-84-0, GenBank A63621
                                                  217893-85-1, GenBank A63622
                                 217893-89-5, GenBank A63627 217893-90-8,
     217893-86-2, GenBank A63624
                     217893-91-9, GenBank A63629 217893-92-0, GenBank A63630
     GenBank A63628
     367319-52-6
                  367319-53-7
                                367319-54-8
                                             367319-55-9
                                                            367319-56-0
     367319-58-2
                   367319-59-3
                                367319-60-6
                                              367319-61-7
                                                            367319-62-8
     367319-63-9
                  367319-64-0
                                367319-65-1
                                              367319-66-2
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     229477-44-5
                   244008-03-5, PN: WO9947540 SEQID: 3 unclaimed DNA
ΙT
     244008-06-8, PN: WO9947540 SEQID: 4 unclaimed DNA 244008-07-9, PN:
                                       244008-08-0, PN: WO9947540 SEQID: 6
    WO9947540 SEQID: 5 unclaimed DNA
                   244008-09-1, PN: WO9947540 SEQID: 7 unclaimed DNA
     unclaimed DNA
     244008-12-6, 8: PN: WO0183510 SEQID: 8 unclaimed DNA
                                                           244008-13-7, PN:
    WO9947540 SEQID: 9 unclaimed DNA 244008-14-8, PN: WO9947540 SEQID: 10
     unclaimed DNA
                    367273-46-9
                                 367273-47-0
                                                367273-48-1
                                                             370571-84-9
     RL: PRP (Properties).
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
     114949-22-3P, Activin
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (βc; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
    ANSWER 9 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     2001:781077 HCAPLUS
DN
     135:348849
ED
     Entered STN: 26 Oct 2001
TΙ
    Albumin fusion proteins with therapeutic proteins for
     improved shelf-life
    Rosen, Craig A.; Haseltine, William A.
ΙN
    Human Genome Sciences, Inc., USA
PΑ
SO
     PCT Int. Appl., 413 pp.
     CODEN: PIXXD2
DΤ
     Patent
    English
LA
IC
     ICM C12N
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 3, 15
FAN.CNT 7
     PATENT NO.
                     KIND DATE
                                          APPLICATION NO.
                                                           DATE
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                                          WO 2001-US11850 20010412
PΙ
     WO 2001079442
                      Α2
                            20011025
                     A3
     WO 2001079442
                            20020606
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,

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DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                           AU ·2001-64563
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                       Α5
                            20011030
                                                             20010412
     EP 1276849
                       Α2
                            20030122
                                           EP 2001-938994
                                                             20010412
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
     US 2003125247
                       Α1
                            20030703
                                           US 2001-833041
                                                             20010412
     US 2003171267
                            20030911
                                           US 2001-833117
                                                             20010412
                       Α1
     US 2003199043
                       A1
                            20031023
                                           US 2001-832501
                                                             20010412
     JP 2003531590
                       T2
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                                           JP 2001-577426
                                                             20010412
     US 2003219875
                       A1
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                                           US 2001-833118
                                                             20010412
     US 2004010134
                       Α1
                            20040115
                                           US 2001-833245
                                                             20010412
PRAI US 2000-229358P
                       Ρ
                            20000412
     US 2000-199384P
                       Ρ
                            20000425
     US 2000-256931P
                       Ρ
                            20001221
     WO 2001-US11850
                       W
                            20010412
AB
     The present invention encompasses fusion proteins of
     albumin with various therapeutic proteins, and in particular
     various antibodies. Therapeutic proteins may be stabilized to extend the
     shelf-life, and/or to retain the therapeutic protein's
     activity for extended periods of time in solution, in vitro and/or in vivo,
     by genetically or chemical fusing or conjugating the therapeutic
     protein to albumin or a fragment or variant of albumin
        Use of albumin fusion proteins may also reduce the
     need to formulate the protein solns. with large excesses of carrier
     proteins to prevent loss of therapeutic proteins due to factors such as
     binding to the container. Nucleic acid mols. encoding the albumin
     fusion proteins of the invention are also encompassed by the
     invention, as are vectors containing these nucleic acids, host cells
     transformed with these nucleic acids vectors, and methods of making the
     albumin fusion proteins of the invention and using these
     nucleic acids, vectors, and/or host cells. Thus, plasmid vectors are
     constructed in which DNA encoding the desired therapeutic protein may be
     inserted for expression of the albumin fusion proteins
     in yeast (pPPC0005) and mammalian cells (pC4:HSA). Yeast-derived signal
     sequences from Saccharomyces cerevisiae invertase SUC2 gene, or the
     stanniocalcin or native human serum albumin signal peptides, are
     used for secretion in yeast or mammalian systems, resp. Thus, the
     fusion product of human growth hormone with residues 1-387 of
     human serum albumin retains essentially intact biol. activity
     after 5 wk of incubation in tissue culture media at 37°, whereas
     recombinant human growth hormone used as control lost its biol.
     activity in the first week. Although the potency of the albumin
     fusion proteins is slightly lower than the unfused counterparts in
     rapid bioassays, their biol. stability results in much higher biol.
     activity in the longer term in vitro assay or in vivo assays. Addnl., the
     present invention encompasses pharmaceutical compns. comprising
     albumin fusion proteins and methods of treating,
     preventing, or ameliorating diseases, disorders or conditions using
     albumin fusion proteins of the invention.
ST
     albumin fusion therapeutic protein shelflife
ΙT
     Antigens
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (17-1A, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (B7.2, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CA125, antibodies to; albumin fusion proteins with
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therapeutic proteins for improved shelf-life)

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ΙT
    CD antigens
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CD147, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CD33, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CD48, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CD52, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (CD6, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
    Immunoglobulins
ΙT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (E, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
    Histocompatibility antigens
ΙT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (HLA-DR, antibodies to; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
TT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (HM1.24, antibodies to; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
ΙT
    Cell adhesion molecules
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (ICAM-1 (intercellular adhesion mol. 1), antibodies to; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Immunoglobulin receptors
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (IgG type I, antibodies to; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
ΙT
    Selectins
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (L-, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Integrins
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (LPAM-1 (lymphocyte Peyer's patch high endothelial venule adhesion mol.
        1), antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Blood-group substances
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (Lex, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Blood-group substances
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (Ley, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Immunoglobulins
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (M, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
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ΙT

Histocompatibility antigens

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ΙT

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ΙT

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IT

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Angiogenic factors

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RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (MHC (major histocompatibility complex), class I, antibodies to;
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
Histocompatibility antigens
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (MHC (major histocompatibility complex), class II, antibodies to;
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
Proteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (NogoA, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (Nsf2, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Glycoproteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (P170, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Cell adhesion molecules
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (SC-1, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Proteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (SF-25, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Antigens
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (SSEA-1 (stage-specific embryonic antigen 1), antibodies to;
   albumin fusion proteins with therapeutic proteins for
   improved shelf-life)
Antigens
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (TAG-72 (tumor-associated glycoprotein 72), antibodies to; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Cell adhesion molecules
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (VCAM-1, antibodies to; albumin fusion proteins
   with therapeutic proteins for improved shelf-life)
Drug delivery systems
Gene therapy
Molecular cloning
   (albumin fusion proteins with therapeutic proteins
   for improved shelf-life)
Antibodies
Cell adhesion molecules
Cytokines
Enzymes, biological studies
  Fusion proteins (chimeric proteins)
Growth factors, animal
Immunoglobulins
  Interferons
Synthetic gene
Tumor necrosis factor receptors
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (albumin fusion proteins with therapeutic proteins
   for improved shelf-life)
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ΙT

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CD14 (antigen)
CD2 (antigen)
CD20 (antigen)
CD22 (antigen)
CD3 (antigen)
CD30 (antigen)
CD38 (antigen)
CD4 (antigen)
CD40 (antigen)
CD44 (antigen)
CD45 (antigen)
CD5 (antigen)
CD8 (antigen)
CD80 (antigen)
CD80 (antigen)
CTLA-4 (antigen)
Carcinoembryonic antigen
Epidermal growth factor receptors
Fas antigen
Integrins
Interleukin 4 receptors
Interleukin 5
LFA-1 (antigen)
Mucins
TCR (T cell receptors)
Transferrin receptors
neu (receptor)
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Mucins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (episialins, antibodies to; albumin fusion proteins
   with therapeutic proteins for improved shelf-life)
Signal peptides
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
   (for improved secretion in yeast or mammalian cells; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Glycoproteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (gD, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Envelope proteins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (gp120env, antibodies to; albumin fusion proteins
   with therapeutic proteins for improved shelf-life)
Glycoproteins, specific or class
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (gpII, antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Animal cell
   (mammalian, recombinant expression host; albumin
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Agglutinins and Lectins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (mannan-binding, antibodies to; albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Antibodies
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
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use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (monoclonal; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
ΙT
    Plasmid vectors
        (pScCHSa, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
ΙT
     Interleukin 6 receptors
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (receptor-associated glycoprotein gp130, antibodies to; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
TΤ
     Saccharomyces cerevisiae
    Yeast
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
ΙT
    Albumins, biological studies
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (serum; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
TT
    Genetic element
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (signal sequence, for improved secretion in yeast or mammalian cells;
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
    Antibodies
IT
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (single chain; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Venoms
        (snake, antibodies to; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (therapeutic; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Globulins, biological studies
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (thymocyte, antibodies to; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
ΙT
    Antigens
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (tumor-associated, antibodies to; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Interleukin 2 receptors
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unclaimed DNA

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RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\alpha-chain, antibodies to; albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Interferons
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (α ; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
Integrins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\alpha IIb\beta 3, antibodies to;
                            albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Vitronectin receptors
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\alpha v \beta 3, antibodies to;
                           albumin fusion
   proteins with therapeutic proteins for improved shelf-
   ·life)
Integrins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\alpha 4\beta 1, \text{ antibodies to;}
                           albumin fusion
   proteins with therapeutic proteins for improved shelf-
   life)
Chemokine receptors
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\beta \text{ chemokine receptor CCR5, antibodies to; albumin})
   fusion proteins with therapeutic proteins for improved
   shelf-life)
Integrins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (\beta 2, \text{ antibodies to; albumin fusion proteins})
   with therapeutic proteins for improved shelf-life)
9002-67-9P, Luteinizing hormone
                                   9002-68-0P, FSH
                                                      9002-72-6P, Growth
          9004-10-8P, Insulin, biological studies
                                                      11096-26-7P,
                 67763-96-6P, Insulin-like growth factor 1
Erythropoietin
                                                              83869-56-1P,
         143011-72-7P, G-CSF
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (albumin fusion proteins with therapeutic proteins
   for improved shelf-life)
156586-89-9
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (albumin fusion proteins with therapeutic proteins
   for improved shelf-life)
11016-39-0, Properdin
                        19600-01-2, Ganglioside GM2
                                                        20830-75-5, Digoxin
99085-47-9, CD55 antigen
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (antibodies to; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
127361-02-8DP, Albumin (human blood serum clone HSA-II/HSA-I-A
protein moiety reduced), full-length or subfragment fusion
RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
use); BIOL (Biological study); PREP (Preparation); USES (Uses)
   (nucleotide sequence; albumin fusion proteins with
   therapeutic proteins for improved shelf-life)
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              167731-74-0, PN: US5962255 SEQID: 56 unclaimed DNA
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US5962255 SEQID: 58 unclaimed DNA
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167731-78-4, PN: US5962255 SEQID: 61 unclaimed DNA

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167731-81-9 167732-10-7
                  167731-80-8
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    GenBank A63616
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    367319-59-3
    367319-64-0
                  367319-65-1
                                367319-66-2
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
       proteins with therapeutic proteins for improved shelf-
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    WO9947540 SEQID: 6 unclaimed DNA
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                    244008-12-6, 8: PN: WO0183510 SEQID: 8 unclaimed DNA
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    WO9947540 SEQID: 10 unclaimed DNA
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    367273-48-1
    RL: PRP (Properties)
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
    ANSWER 10 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
    2001:780938 HCAPLUS
    135:322686
    Entered STN: 26 Oct 2001
    Albumin fusion proteins with therapeutic proteins for
    improved shelf-life
    Rosen, Craig A.; Sadeghi, Homayoun; Prior, Christopher P.;
    Turner, Andrew John
    Human Genome Sciences, Inc., USA; Principia Pharmaceutical
    Corporation
    PCT Int. Appl., 328 pp.
    CODEN: PIXXD2
    Patent
    English
    ICM C07K001-00
    ICS A01N037-18
     63-3 (Pharmaceuticals)
    Section cross-reference(s): 3, 15
FAN.CNT 7
                                          APPLICATION NO. DATE
    PATENT NO.
                     KIND DATE
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                                                           _____
    WO 2001079258
                      A1
                           20011025
                                          WO 2001-US12008 20010412
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            CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
            HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
            LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
            RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
            VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
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Antidiabetic agents

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20030115
                                           EP 2001-932549
                                                            20010412
     EP 1274720
                       Α1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                           US 2001-833041
                                                            20010412
     US 2003125247
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                            20030703
                            20030911
                                           US 2001-833117
                                                            20010412
     US 2003171267
                       Α1
     JP 2003530838
                       T2
                            20031021
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                                                            20010412
                            20031023
                                           US 2001-832501
    US 2003199043
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                                                            20010412
                                           US 2001-833118
     US 2003219875
                       Α1
                            20031127
                                                            20010412
                                           US 2001-833245
     US 2004010134
                       Α1
                            20040115
                                                            20010412
PRAI US 2000-229358P
                       Ρ
                            20000412
     US 2000-199384P
                      Ρ
                            20000425
     US 2000-256931P
                      Ρ
                            20001221
    WO 2001-US12008 W
                            20010412
     The present invention encompasses fusion proteins of
AΒ
     albumin with various therapeutic proteins, and in particular, with
     interleukin 2, calcitonin, growth hormone-releasing factor,
     interferon \beta , parathyroid hormine, and insulin-like
     growth factor 1. Therapeutic proteins may be stabilized to extend the
     shelf-life, and/or to retain the therapeutic protein's
     activity for extended periods of time in solution, in vitro and/or in vivo,
     by genetically or chemical fusing or conjugating the therapeutic
    protein to albumin or a fragment or variant of albumin
        Use of albumin fusion proteins may also reduce the
     need to formulate the protein solns. with large excesses of carrier
    proteins to prevent loss of therapeutic proteins due to factors such as
     binding to the container. Nucleic acid mols. encoding the albumin
     fusion proteins of the invention are also encompassed by the
     invention, as are vectors containing these nucleic acids, host cells
     transformed with these nucleic acids vectors, and methods of making the
     albumin fusion proteins of the invention and using these
     nucleic acids, vectors, and/or host cells. Thus, plasmid vectors are
     constructed in which DNA encoding the desired therapeutic protein may be
     inserted for expression of the albumin fusion proteins
     in yeast (pPPC0005) and mammalian cells (pC4:HSA).
                                                         Yeast-derived signal
     sequences from Saccharomyces cerevisiae invertase SUC2 gene, or the
     stanniocalcin or native human serum albumin signal peptides, are
     used for secretion in yeast or mammalian systems, resp. Thus, the
     fusion product of human growth hormone with residues 1-387 of
     human serum albumin retains essentially intact biol. activity
     after 5 wk of incubation in tissue culture media at 37°, whereas
     recombinant human growth hormone used as control lost its biol.
     activity in the first week. Although the potency of the albumin
     fusion proteins is slightly lower than the unfused counterparts in
     rapid bioassays, their biol. stability results in much higher biol.
     activity in the longer term in vitro assay or in vivo assays. Addnl., the
     present invention encompasses pharmaceutical compns. comprising
     albumin fusion proteins and methods of treating,
     preventing, or ameliorating diseases, disorders or conditions using
     albumin fusion proteins of the invention.
ST
     albumin fusion therapeutic protein shelflife
ΙT
     Hepatitis
        (C, agents for treatment of; albumin fusion
        proteins with therapeutic proteins for improved shelf-
     Antitumor agents
ΙT
        (Kaposi's sarcoma; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
     Antitumor agents
        (acute myelogenous leukemia; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     Anti-AIDS agents
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المراجعة

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Antirheumatic agents
     Drug delivery systems
    Gene therapy
     Immunosuppressants
    Molecular cloning
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
ΙT
    Cell adhesion molecules
     Cytokines
    Enzymes, biological studies
      Fusion proteins (chimeric proteins)
    Growth factors, animal
       Interferons
     Interleukin 2
     Synthetic gene
     Tumor necrosis factor receptors
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     Signal peptides
IT
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (for improved secretion in yeast or mammalian cells; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
     Intestine, disease
ΙT
        (inflammatory, agents for treatment of; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Kidney, neoplasm
     Lung, neoplasm
     Ovary, neoplasm
        (inhibitors; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Antitumor agents
        (kidney; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Antitumor agents
        (leukemia; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Antitumor agents
        (lung; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Animal cell
        (mammalian, recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
     Antitumor agents
ΙT
        (melanoma, metastasis; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Antitumor agents
        (melanoma; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
ΙT
     Antitumor agents
        (non-Hodgkin's lymphoma; albumin fusion proteins
        with therapeutic proteins for improved shelf-life)
ΙT
     Antitumor agents
        (ovary; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
     Plasmid vectors
        (pC4:HSA, for mammalian cell expression; albumin
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fusion proteins with therapeutic proteins for improved

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shelf-life)
IT
    Plasmid vectors
        (pPPC0005, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Plasmid vectors
IT
        (pScCHSa, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Plasmid vectors
        (pScNHSA, for yeast expression; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
    Saccharomyces cerevisiae
IT
    Yeast
        (recombinant expression host; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
    Kidney, neoplasm
IT
        (renal-cell carcinoma, metastasis, inhibitors; albumin
        fusion proteins with therapeutic proteins for improved
        shelf-life)
IT
    Antitumor agents
        (renal-cell carcinoma, metastasis; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
IT
    Albumins, biological studies
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (serum; albumin fusion proteins with therapeutic
        proteins for improved shelf-life)
IT
    Genetic element
    RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (signal sequence, for improved secretion in yeast or mammalian cells;
        albumin fusion proteins with therapeutic proteins for
        improved shelf-life)
IT
    Antibodies
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (single chain; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Multiple sclerosis
        (therapeutic agents; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
     Proteins, specific or class
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (therapeutic; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
IT
    Interferons
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (\alpha; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
ΙT
    Interferons
    RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
    use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (\beta ; albumin fusion proteins with
       therapeutic proteins for improved shelf-life)
     9002-64-6P, Parathyroid hormone 9002-67-9P, Luteinizing hormone
ΙT
     9002-68-0P, FSH
                     9002-72-6P, Growth hormone
                                                    9004-10-8P, Insulin,
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9007-12-9P, Calcitonin

9034-39-3P, Growth

biological studies

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63-3 (Pharmaceuticals)

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11096-26-7P, Erythropoietin
     hormone-releasing factor
                                                               67763-96-6P,
     Insulin-like growth factor 1
                                   83869-56-1P, GM-CSF
                                                          143011-72-7P, G-CSF
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (albumin fusion proteins with therapeutic proteins
        for improved shelf-life)
     127361-02-8DP, Albumin (human blood serum clone HSA-II/HSA-I-A
     protein moiety reduced), full-length or subfragment fusion
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (nucleotide sequence; albumin fusion proteins with
       therapeutic proteins for improved shelf-life)
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     RL: PRP (Properties)
        (unclaimed nucleotide sequence; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     367510-76-7
     RL: PRP (Properties)
        (unclaimed protein sequence; albumin fusion
        proteins with therapeutic proteins for improved shelf-
        life)
     131748-18-0
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                                               367273-48-1
                   367273-46-9
     RL: PRP (Properties)
        (unclaimed sequence; albumin fusion proteins with
        therapeutic proteins for improved shelf-life)
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Beth Israel Hospital Association; WO 9618412 A1 1996 HCAPLUS
(2) Lee; Pharm Dev Tech 1999, V4(2), P269 HCAPLUS
(3) Rhone-Poulenc Rorer S A; WO 9315199 A1 1993 HCAPLUS
(4) Rhone-Poulenc Rorer S A; WO 9315211 A1 1993 HCAPLUS
(5) Takahashi; Peptides 1997, V18(3), P439 HCAPLUS
(6) Yeh; Prc Nat Acad Sci USA 1992, V69, P1904
     ANSWER 11 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
     2001:763025 HCAPLUS
     135:335111
     Entered STN: 19 Oct 2001
     Albumin fusion proteins with therapeutic proteins for improved shelf-life
     Rosen, Craig A.; Haseltine, William A.
     Human Genome Sciences, Inc., USA
     PCT Int. Appl., 2102 pp.
     CODEN: PIXXD2
     Patent
     English
     ICM C07H021-04
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Section cross-reference(s): 3, 15

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FAN.CNT 7
                                          APPLICATION NO. DATE
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                                    WO 2001-US11988 20010412
     WO 2001077137
                     A1
                           20011018
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             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
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             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                      A1
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                                          EP 2001-944114 20010412
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
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AB
     The present invention encompasses fusion proteins of albumin with various
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therapeutic proteins. Therapeutic proteins may be stabilized to extend the shelf-life, and/or to retain the therapeutic protein's activity for extended periods of time in solution, in vitro and/or in vivo, by genetically or chemical fusing or conjugating the therapeutic protein to albumin or a fragment or variant of albumin. Use of albumin fusion proteins may also reduce the need to formulate the protein solns. with large excesses of carrier proteins to prevent loss of therapeutic proteins due to factors such as binding to the container. Nucleic acid mols. encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Thus, plasmid vectors are constructed in which DNA encoding the desired therapeutic protein may be inserted for expression of the albumin fusion proteins in yeast (pPPC0005) and mammalian cells (pC4:HSA). Yeast-derived signal sequences from Saccharomyces cerevisiae invertase SUC2 gene, or the stanniocalcin or native human serum albumin signal peptides, are used for secretion in yeast or mammalian systems, resp. Thus, the fusion product of human growth hormone with residues 1-387 of human serum albumin retains essentially intact biol. activity after 5 wk of incubation in tissue culture media at 37°, whereas recombinant human growth hormone used as control lost its biol. activity in the first week. Although the potency of the albumin fusion proteins is slightly lower than the unfused counterparts in rapid bioassays, their biol. stability results in much higher biol. activity in the longer term in vitro assay or in vivo assays. Addnl., the present invention encompasses pharmaceutical compns. comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disorders or conditions using albumin fusion proteins of the invention.

ST albumin fusion therapeutic protein shelflife

IT Drug delivery systems

Gene therapy

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+7

Molecular cloning

(albumin fusion proteins with therapeutic proteins for improved shelf-life)

IT Cell adhesion molecules

و والمترام

ورونية

IT

Cytokines Enzymes, biological studies Fusion proteins (chimeric proteins) Growth factors, animal Interferons Synthetic gene Tumor necrosis factor receptors RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (albumin fusion proteins with therapeutic proteins for improved shelf-life) IT Signal peptides RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (for improved secretion in yeast or mammalian cells; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT (mammalian, recombinant expression host; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT Plasmid vectors (pC4:HSA, for mammalian cell expression; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT Plasmid vectors (pPPC0005, for yeast expression; albumin fusion proteins with therapeutic proteins for improved shelf-life) Plasmid vectors IΤ (pScCHSa, for yeast expression; albumin fusion proteins with therapeutic proteins for improved shelf-life) IT (pScNHSA, for yeast expression; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT Saccharomyces cerevisiae Yeast (recombinant expression host; albumin fusion proteins with therapeutic proteins for improved shelf-life) Albumins, biological studies IΤ RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (serum; albumin fusion proteins with therapeutic proteins for improved shelf-life) TT ' Genetic element RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (signal sequence, for improved secretion in yeast or mammalian cells; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙT Antibodies RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (single chain; albumin fusion proteins with therapeutic proteins for improved shelf-life) ΙΤ Proteins, specific or class RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (therapeutic; albumin fusion proteins with therapeutic proteins for improved shelf-life) IT' Interferons RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)  $(\alpha;$  albumin fusion proteins with therapeutic proteins for improved shelf-life)

9002-68-0P, FSH

9004-10-8P, Insulin, biological studies

9002-72-6P, Growth

11096-26-7P,

9002-67-9P, Luteinizing hormone

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67763-96-6P, Insulin-like growth factor 1
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             143011-72-7P, G-CSF
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        (albumin fusion proteins with therapeutic proteins for improved
        shelf-life)
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(unclaimed sequence; albumin fusion proteins with therapeutic proteins for improved shelf-life)

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THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 3
(1) Delta Biotechnology Limited; EP 0322094 A1 1989 HCAPLUS
(2) Delta Biotechnology Limited; WO 9724445 A1 1997 HCAPLUS
(3) Human Genome Sciences Inc; WO 9734997 A1 1997 HCAPLUS
L66
    ANSWER 12 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
     2000:609058 HCAPLUS
ΑN
DN
     133:168425
     Entered STN: 01 Sep 2000
ED
     Suppository of recombinant human interferon .
ΤI
     alpha.2a
     Chen, Weijia; Zheng, Hui; Zhang, Yan; Wang, Dongqian
ΙN
     Changchun Biological Product Inst., Ministry of Public Health, Peop. Rep.
PΑ
SO
     Faming Zhuanli Shenqing Gongkai Shuomingshu, 5 pp.
     CODEN: CNXXEV
DT
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LA
     Chinese
     ICM A61K009-02
IC
     ICS A61K038-21
     63-6 (Pharmaceuticals)
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PRAI CN 1999-105589
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     Suppository of interferon \alpha 2a comprise
     recombinant human interferon \alpha 2a solution
     (0.5 MIU per suppository) 14, glycerol 58, gelatin 26, and human serum
     albumin 2%. The preparation process involves mixing glycerol with
     gelatin, standing overnight, sterilizing for 20-30 min, cooling to
     40-56Φ', adding recombinant human interferon .
     alpha.2a, and shaping.
     recombinant human interferon alpha 2a
ST
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     Albumins, biological studies
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        interferon \alpha 2a)
ΙT
     Anti-inflammatory agents
     Antitumor agents
     Antiviral agents
     Skin, disease
        (suppository of recombinant human interferon
        \alpha 2a)
IT
     Gelatins, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
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ΙT
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        \alpha 2a)
     56-81-5, Glycerol, biological studies
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وبالمواكنة

ور وأنتقه

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    ANSWER 13 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
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     Entered STN: 10 Dec 1999
     Recombinant human interferon \beta -1A (
TI
     IFN-beta-1A) formulation
IN
     Alam, John; Rogge, Mark; Goelz, Susan
    Biogen, Inc., USA
PΑ
SO
     PCT Int. Appl., 28 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM A61K038-21
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 15
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     EP 1082132
                     A1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
                                           JP 2000-551797
                                                            19980529 <--
                     Т2
     JP 2002516874
                            20020611
                                           EE 2000-20000069419980529 <--
     EE 200000694
                       Α
                            20020617
                                           NO 2000-6022
                                                            20001128 <--
                            20010126
     NO 2000006022
                      Α
PRAI WO 1998-US7242
                     Α
                            19980529 <--
     Liquid compns. comprising a buffer of pH about 7.2, recombinant
     interferon-\beta and 15 mg/mL of human serum
     albumin, and kits for parenteral administration comprising said
     compns. are disclosed.
     recombinant interferon beta formulation
ST
IT
     Medical goods
        (alc. swabs; recombinant human interferon
        \beta -1A (IFN-beta-1A) formulation)
ΙT
     Medical goods
        (bandages, adhesive; recombinant human interferon
        \beta -1A (IFN-beta-1A) formulation)
ΙT
     Buffers
     Molecular cloning
     Needles (tools)
     Syringes
     рΗ
        (recombinant human interferon \beta -1A (
        IFN-beta-1A) formulation)
     Albumins, biological studies
IT
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL: (Biological study); PROC (Process); USES (Uses)
        (serum, human; recombinant human interferon
        \beta -1A (IFN-beta-1A) formulation)
     Interferons
     RL: BPN (Biosynthetic preparation); PEP (Physical, engineering or chemical
     process); THU (Therapeutic use); BIOL (Biological study); PREP
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(Preparation); PROC (Process); USES (Uses)
        (β ; recombinant human interferon
        β -1A (IFN-beta-1A) formulation)
     145258-61-3, Interferon \beta 1 (human fibroblast
ΙT
     protein moiety)
     RL: PEP (Physical, engineering or chemical process); THU (Therapeutic
     use); BIOL (Biological study); PROC (Process); USES (Uses)
        (recombinant human interferon β -1A (
        IFN-beta-1A) formulation)
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
(1) Alam, J; Pharmaceutical Research 1997, V14(4), P546 HCAPLUS
(2) Anon; http://www.healthdirect.com/usenew/pressrel/p biogel.htm 1996
(3) Salmon, P; Journal of Interferon and Cytokine Research 1996, V16(10), P759
(4) US Food and Drug Administration-Interferon Beta-1A, Biogen, Inc;
   http://www.fda.gov/cber/products/ifnbbio051796.htm,
   http://www.fda.gov/cber/label/infbbio051796lb.pdf 1998
L66 ANSWER 14 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
     1999:563880 HCAPLUS
ΑN
DN
     131:161626
ΕD
     Entered STN:
                   08 Sep 1999
TΙ
     Oral recombinant human \alpha -interferon
     compositions
     Dong, Yilan; Cheng, Xiaogeng; Lin, Yuxin; Wang, Shiwen; Liu, Zhenhao;
IN
PA
     Changchun Institute of Biological Products, Ministry of Public Health,
     Peop. Rep. China
     Faming Zhuanli Shenging Gongkai Shuomingshu, 8 pp.
SO
     CODEN: CNXXEV
DT
     Patent
LA
     Chinese
IC
     ICM A61K038-21
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1, 15
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                             DATE
     ______
                      ____
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                                            _____
     CN 1116951
                            19960221
                                           CN 1995-101216
                                                             19950125 <--
                       Α
PRAI CN 1995-101216
                            19950125
                                      <--
     Title compns. as antiviral agents contain recombinant human .
     alpha.-interferon 100-500 IU, thymosin F5 isolated from
     calf's thymus gland 1-20 \mug, stabilizers and conventional medical
     additives. The stabilizers are selected from human serum albumin
     , cattle serum albumin, \beta-cyclodextrin and PEG 800.
ST
     recombinant human interferon tablet antiviral
IT
     Antiviral agents
     Stabilizing agents
        (oral recombinant human \alpha -interferon
        compns.)
ΙT
     Polyoxyalkylenes, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (oral recombinant human α -interferon
        compns.)
ΙT
     Drug delivery systems
        (oral; oral recombinant human \alpha -
        interferon compns.)
     Albumins, biological studies
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (serum, human or bovine; oral recombinant human
        \alpha -interferon compns.)
     Drug delivery systems
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(tablets; oral recombinant human \alpha -
        interferon compns.)
IT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (α , recombinant human; oral
        recombinant human α -interferon
        compns.)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (α -2a, recombinant human; oral
        recombinant human \alpha - interferon
        compns.)
TΤ
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\alpha - 2b, recombinant human; oral
        recombinant human α -interferon
        compns.)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (α 1, recombinant human; oral
        recombinant human α -interferon
        compns.)
IT
     61512-21-8, Thymosin
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (F5; oral recombinant human \alpha -
        interferon compns.)
     7585-39-9, \beta-Cyclodextrin
                                  25322-68-3
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (oral recombinant human \alpha -interferon
        compns.)
    ANSWER 15 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
     1997:756962 HCAPLUS
AN
DN
     128:16442
     Entered STN: 04 Dec 1997
ED
     Stabilization of interferons in aqueous solution for manufacture
TI
     of sublingually administered tablets
     Rothschild, Peter R.
ΙN
     Feronpatent Limited, Ire.; Rothschild, Peter R.
PΑ
SO
     PCT Int. Appl., 12 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM A61K038-21
IC
     ICS A61K009-20
     63-6 (Pharmaceuticals)
FAN.CNT 1
                      KIND DATE
                                           APPLICATION NO.
                                                             DATE
     PATENT NO.
                                            _____
     ______
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                            _____
                                                             _____
                                                             19970509 <--
                            19971113
                                           WO 1997-IB531
PΙ
     WO 9741885
                       A1
             AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
             DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,
             PT, RO, RU, SD, SE, SG, SI, SK, TJ, TM, TR, TT, UA, UG, US, UZ,
             VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB,
             GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN,
             ML, MR, NE, SN, TD, TG
                                            AU 1997-24011
                                                             19970509 <--
     AU 9724011
                            19971126
                       Α1
                                            EP 1997-919596
                                                             19970509 <--
                            19990609
     EP 920329
                       Α1
                            20020925
     EP 920329
                       В1
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI
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E
                                           AT 1997-919596
                                                             19970509 <--
    AT 224725
                            20021015
                       Т3
                                           ES 1997-919596
                                                             19970509 <--
                            20030401
     ES 2184084
                            19960509
PRAI WO 1996-IB433
                       Α
                                      <--
     WO 1997-IB531
                       W
                            19970509
                                      <--
AB
    Natural and recombinant interferons are stabilized
    with bidistd. water, lactose, albumin, sodium mono- and
     dihydrogen phosphates, (C5H10O5)n, such as arabic gum, dissolved and diluted
     in 20 % ethanol solution to the fourth decimal by homeopathic method. The
     final solution is sprayed on to an excipient comprising of 20 % arabic gum,
     30 % lactose and 50 % starch for manufacturing tablets of 100 mg each
containing 200
     I.U. of human alfa-interferon. The tablets are sublingually
     administered to the patient for treatment of viral infections
     sensitive to interferon. Preparation of sublingual tablets according
     above method is disclosed.
     stabilization interferon polysaccharide sublingual
ST
     pharmaceutical tablet
ΙT
     Hepatitis
        (B; stabilization of interferons in aqueous solution for manufacture of
        sublingually administered tablets)
IT
        (C; stabilization of interferons in aqueous solution for manufacture of
        sublingually administered tablets)
ΙT
        (homeopathy; stabilization of interferons in aqueous solution for
        manufacture of sublingually administered tablets)
ΙT
     Antitumor agents
     Stabilizing agents
        (stabilization of interferons in aqueous solution for manufacture of
        sublingually administered tablets)
     Albumins, biological studies
IT
       Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (stabilization of interferons in aqueous solution for manufacture of
        sublingually administered tablets)
     Drug delivery systems
ΙT
        (tablets, sublingual; stabilization of interferons in aqueous
        solution for manufacture of sublingually administered tablets)
ΙT
        (viral; stabilization of interferons in aqueous solution for manufacture
        of sublingually administered tablets)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\alpha ; stabilization of interferons in aqueous solution
        for manufacture of sublingually administered tablets)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\beta ; stabilization of interferons in aqueous solution
        for manufacture of sublingually administered tablets)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\gamma; stabilization of interferons in aqueous solution for
        manufacture of sublingually administered tablets)
                      7558-79-4, Sodium monohydrogen phosphate
                                                                    7558-80-7,
IT
     63-42-3, Lactose
     Sodium dihydrogen phosphate 9000-01-5, Arabic gum
                                                          9005-25-8, Starch,
     biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (stabilization of interferons in aqueous solution for manufacture of
        sublingually administered tablets)
L66 ANSWER 16 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
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1996:635884 HCAPLUS

125:308823

ΑN

DN

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JP 10500125

T2

19980106

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ΕD
     Entered STN: 28 Oct 1996
     Shelf-life of recombinant human interferon .
ΤI
     alpha.2b under different storage conditions
     Barberia, Daisy; Vega, Maribel; Ferrero, Joel; Duany, Lady; Moya, Galina;
ΑU
     Curras, Tania; Martinez, Maida; Cruz, Asterio; Gil, Miriela; Quintana,
     Marisel
     Centro de Ingenieria Genetica y Biotecnologia, Havana, Cuba
CS
     Biotecnologia Aplicada (1996), 13(3), 190-194
SO
     CODEN: BTAPEP; ISSN: 0864-4551
₽B
     Sociedad Iberolatinoamericana de Biotecnologia Aplicada a la Salud
DT
     Journal
LA
     Spanish
CC.
     63-5 (Pharmaceuticals)
AΒ
     The stability test studies under accelerated and normal storage conditions
     carried out with recombinant human alpha 2b interferon
     (hu-r alpha 2b IFN) in phosphate buffer 0.1M, pH 7.0, with and without
     albumin, in order to establish its shelf-life at refrigerating and
     frozen conditions. According to the accelerated study the authors
     concluded that no alterations will interfere with the recognition of hu-r
     alpha 2b IFN in ELISA in at least five years when stored at -70 or
     -20°. Otherwise, when stored at 4°, a loss of 10% may occur
     in one year. The authors corroborated this when the presence of new
     structures which might affect the protein immunol. recognition were
     detected by RP-HPLC. No stabilizing properties of albumin on
     hu-r alpha 2b IFN were observed at least when it is in phosphate buffer 0.1M,
     pH 7.0 and under accelerated storing conditions.
ST
     interferon stability denaturation freezing
ΙT
     Albumins, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (shelf-life of recombinant human interferon
        α 2b under different storage conditions)
ΙT
     Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\alpha - 2b, shelf-life of recombinant
        human interferon α 2b under
        different storage conditions)
    ANSWER 17 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     1996:43019 HCAPLUS
DN
     124:66661
ΕD
     Entered STN: 23 Jan 1996
     Stabilized \beta -interferon liquid formulations
TΙ
IN
     Samaritani, Fabrizio; Natale, Patrizia
PA
     Applied Research Systems ARS Holding N.V., Neth.
SO
     PCT Int. Appl., 17 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
     ICM A61K038-21
IC
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
     PATENT NO.
                                           APPLICATION NO.
                      KIND DATE
                                                            DATE
     ______
                      ____
                            -----
                                           -----
     WO 9531213
                      A1
                            19951123
                                           WO 1995-EP1825
                                                            19950515 <--
         W: AU, CA, JP, US
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     CA 2190465
                      AA
                            19951123
                                           CA 1995-2190465 19950515 <--
     AU 9526704
                            19951205
                                           AU 1995-26704
                                                            19950515 <--
                       Α1
     AU 704827
                       B2
                            19990506
     EP 759775
                       Α1
                            19970305
                                           EP 1995-921749
                                                            19950515 <--
                       В1
                            20000726
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R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE

JP 1995-529360 19950515 <--

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AT 194917
                            2000081.5
                                            AT 1995-921749
                                                             19950515 <--
                       Т3
                                            ES 1995-921749
                                                             19950515 <--
     ES 2148526
                            20001016
PRAI IT 1994-RM300
                       Α
                            19940516
                                      <--
    WO 1995-EP1825
                       W
                            19950515
                                      <--
     \beta -Interferon liquid formulations are stabilized
AB
     with a polyol, a nonreducing sugar, or an amino acid. In particular, the
     formulations are stabilized with a polyol, such as mannitol. The
     formulations, preferably, furthermore comprise a buffer, such as acetate
     buffer at a pH 3-4 and human albumin at a min. quantity. The .
    beta.-interferon is preferably recombinant.
     interferon soln stabilizer polyol albumin buffer;
ST
    mannitol albumin acetate buffer interferon stability
IT
     Buffer substances and systems
        (acetate; stabilized \beta -interferon liquid
        formulations)
ΙT
     Albumins, biological studies
     Amino acids, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (stabilized \beta -interferon liquid formulations)
     Carbohydrates and Sugars, biological studies
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (nonreducing, stabilized \beta -interferon liquid
        formulations)
IT
     Alcohols, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (polyhydric, stabilized \beta -interferon liquid
        formulations)
IT
     Pharmaceutical dosage forms
        (solns., stabilized \beta -interferon liquid
        formulations)
     Interferons
TΤ
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (\beta , recombinant; stabilized \beta -
        interferon liquid formulations)
     56-40-6, Glycine, biological studies
                                            57-50-1, Saccharose, biological
ΙT
              69-65-8, D-Mannitol
     studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (stabilized \beta -interferon liquid formulations)
    ANSWER 18 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
1.66
ΑN
     1995:498838 HCAPLUS
DN
     122:248213
     Entered STN: 20 Apr 1995
ΕD
     Influence of human serum albumin content in
TΤ
     formulations on the bioequivalency of interferon alfa-2a given
     by subcutaneous injection in healthy male volunteers
     Zhi, Jianquo; Teller, Stuart B.; Satoh, Hiroko; Koss-Twardy, Susan G.;
ΑU
     Luke, David R.
     Department of Clinical Pharmacokinetics, Hoffmann-La Roche, Inc., Nutley,
CS
     NJ, 07110-1199, USA
     Journal of Clinical Pharmacology (1995), 35(3), 281-4
SO
     CODEN: JCPCBR; ISSN: 0091-2700
DT
     Journal
     English
T.A
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 1
     To determine the influence of human serum albumin (HSA)
AB
     content in formulations on the bioequivalency of recombinant
     interferon \alpha 2a, a double-blind, randomized,
     two-way crossover study was conducted in 24 healthy male volunteers.
     Subjects received a single s.c. injection of 18 million IU of Roferon-A
     reconstituted with either the diluent containing 10 mg of HSA or the HSA-free
     diluent; final HSA contents in the 2 formulations were 15 and 5 mg, resp.
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Administration of the 2 formulations resulted in similar 48-h Roferon-A serum concentration-time profiles and comparable frequency and intensity of adverse events. The statistical anal. using the two one-sided tests procedure showed that both formulations were bioequivalent for pharmacokinetic parameters such as Cmax, tmax, AUC48, and AUC. threefold change in HSA content in formulations does not alter the bioequivalency of Roferon-A. interferon bioavailability bioequivalence injection albumin Drug bioavailability (human serum albumin effect on bioequivalence of recombinant interferon  $\alpha$  2a from s.c. injection in humans) Albumins, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (human serum albumin effect on bioequivalence of recombinant interferon  $\alpha$  2a from s.c. injection in humans) Pharmaceutical dosage forms (injections, s.c., human serum albumin effect on bioequivalence of recombinant interferon  $\alpha$  2a from s.c. injection in humans) Interferons RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)  $(\alpha$  -2a, human serum albumin effect on bioequivalence of recombinant interferon  $\alpha$  2a from s.c. injection in humans) L66 ANSWER 19 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN **1994:6892** HCAPLUS 120:6892 Entered STN: 08 Jan 1994 Novel recombinant human IFN- $\beta$  , its preparation, and pharmaceutical compositions containing it Siklosi, Thomas; Joester, Karl-eduard; Hofer, Hans BIOFERON Biochemische Substanzen GmbH und Co, Germany Eur. Pat. Appl., 19 pp. CODEN: EPXXDW Patent German ICM C07K015-26 ICS C07K003-28; A61K037-66 16-2 (Fermentation and Bioindustrial Chemistry) Section cross-reference(s): 15 FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE -----\_\_\_\_\_ ----EP 529300 A1 19930303 EP 1992-112427 19920721 <--EP 529300 B1 19981014 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, MC, NL, PT, SE DE 4128319 A1 19930304 DE 1991-4128319 19910827 <--AT 1992-112427 AT 172206 F. 19981015 19920721 <--ES 2121804 Т3 ES 1992-112427 19981216 19920721 <--PRAI DE 1991-4128319 19910827 <--A recombinant human  $\beta$  -interferon ( IFN- $\beta$ ) produced in mammalian cells, whose oligosaccharide component comprises biantennary ≥60%, triantennary ≥15%, and tetraantennary 0-5% and contains fucose and ≥80% sialic acid, is useful for treatment of tumors, especially Kaposi's sarcoma. Thus, recombinant IFN- $\beta$  was produced in transfected CHO BIC 8622 cells in MEM containing fetal calf serum and secreted

into the medium in a yield of 1 + 105-1 + 106 IU/L. The

و من المنظوم

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IFN-β was purified by liquid-liquid extraction in a PEG
     2000-salt solution system, affinity chromatog. on Blue Dextran FF, metal
     chelate chromatog. on a Zn2+-loaded chelating Sepharose column, and size
     exclusion chromatog. on Sephacryl. The product showed a purity of >99\% and high stability at -20, +15, or +25\% when mixed with buffered
     human serum albumin and stored for 1-4 wk. Enzymic removal of
     terminal sialic acid residues diminished the stability.
     recombinant beta interferon purifn
ST
     Polyoxyalkylenes, biological studies
ΙT
     Salts, biological studies
     RL: BIOL (Biological study)
         (in \beta -interferon purification, by partition)
ΙT
     Oligosaccharides
     Sialic acids
     RL: BIOL (Biological study)
         (of recombinant \beta -interferon)
ΙT
     Chromatography, gel
         (of β -interferon)
ΙT
     Partition
         (of \beta -interferon, in polyalkylene
        glycol/dextran and polyalkylene glycol/salt systems)
ΙT
     Neoplasm inhibitors
         (recombinant \beta -interferon)
ΙT
     Dyes
         (β -interferon affinity chromatog. on)
     Animal cell line
ΙT
         (CHO, recombinant β -interferon
        manufacture with)
IT
     Neoplasm inhibitors
         (Kaposi's sarcoma, recombinant \beta -
        interferon as)
ΙT
     Chromatography, column and liquid
         (affinity, of \beta -interferon, on dye)
ΙT
     Coordination compounds
     RL: BIOL (Biological study)
         (chelates, stationary phases containing, for \beta -
        interferon chromatog.)
IT
     Interferons
     RL: BIOL (Biological study)
         (\beta , purification of recombinant, for Kaposi's
        sarcoma treatment)
                                                      57-55-6, 1,2-Propanediol,
                   148498-83-3, Blue Sepharose FF
IT
     12236-82-7
             107-21-1, 1,2-Ethanediol, uses
     RL: BIOL (Biological study)
         (in \beta -interferon purification, by affinity
        chromatog.)
                                                              288-32-4, Imidazole, ·
                                71-00-1, Histidine, uses
ΙT
     56-40-6, Glycine, uses
     RL: USES (Uses)
         (in \beta -interferon purification, by metal chelate
         chromatog.)
                                                              25322-68-3,
IT
     62-76-0, Sodium oxalate
                                 68-04-2, Sodium citrate
     Polyethylene glycol 25322-69-4, Polypropylene glycol 7447-40-7, Potassium chloride (KCl), uses 7447-41-8, Lithium chloride, uses
                                                                   7447-40-7,
                                        7558-80-7, Sodium dihydrogen phosphate
     7558-79-4, Disodium phosphate
     7647-14-5, Sodium chloride, uses 7681-11-0, Potassium iodide, uses
                                         7757-82-6, Sodium sulfate, uses
     7681-82-5, Sodium iodide, uses
     7758-11-4, Dipotassium phosphate 7778-80-5, Potassium sulfate, uses
     7783-20-2, Ammonium sulfate, uses 9004-54-0, Dextran, uses
                                                                        12125-02-9,
     Ammonium chloride, uses
     RL: BIOL (Biological study)
         (in \beta -interferon purification, by partition)
     131-48-6, N-Acetylneuraminic acid 1113-83-3 2438-80-4, Fucose
ΙT
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response to)

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83412-55-9
     32181-59-2, N-Acetyllactosamine
                                        78392-81-1
                                                                  84813-89-8
                                148553-76-8 148553-77-9
     123618-73-5
                   131432-29-6
                                                              148553-78-0
     148553-79-1
                                 148553-81-5
                                                148614-65-7
                   148553-80-4
                                                              148615-15-0
     RL: BIOL (Biological study)
        (of recombinant \beta -interferon)
IT
     7440-02-0D, Nickel, chelates 7440-48-4D, Cobalt, chelates
                                                                    7440-50-8D.
     Copper, chelates
                       7440-66-6D, Zinc, chelates 12774-36-6, Sephadex G150
     97599-42-3, Superose 12
                                119332-87-5, Sephacryl S 200 High Resolution
     148499-25-6, TSK-SW 3000
     RL: BIOL (Biological study)
        (\beta -interferon purification by chromatog. on)
     ANSWER 20 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
     1992:468225 HCAPLUS
ΑN
DN
     117:68225
ED
     Entered STN:
                   23 Aug 1992
ΤI
     Human \beta -interferon incubated with muscle
     homogenate is protected by albumin but not by proteinase
ΑU
     Paulesu, L.; Pessina, G. P.; Bocci, V.
     Inst. Gen. Physiol., Univ. Siena, Siena, 53100, Italy
CS
SO
     Proceedings of the Society for Experimental Biology and Medicine (
     1992), 200(3), 414-17
     CODEN: PSEBAA; ISSN: 0037-9727
DT
     Journal
LA
     English
CC
     15-5 (Immunochemistry)
     Section cross-reference(s): 1
AB
     The scarce bioavailability of \beta -interferon (
     IFN-β ) after i.m. administration is probably due
     either to the binding of IFN-\beta to the
     interstitial matrix, or to lymphatic absorption and/or to local breakdown
     by lysosomal proteinases from muscle. In this work, the authors first
     showed that after i.m. injection, the apparent bioavailability of natural
     human IFN-\beta is about 10% of that of
     recombinant IFN-\alpha 2 and then they
     evaluated the effects of proteinase inhibitors and albumin on
     IFN-β incubated at 37° with muscle
     homogenate. IFN biol. activity decreased spontaneously by about 20% after
     incubation for 6 h at 37° in Hanks' solution, but it was almost
     completely lost after incubation with muscle homogenate. Proteinase
     inhibitors (\alpha1-antitrypsin, \alpha2-macroglobulin, aprotinin,
     soybean trypsin inhibitor, leupeptin, EP-459, and EP-475) failed to block
     the inactivation of IFN-\beta by muscle proteinases,
     whereas albumin exerted a partial but consistent protection.
ST
     interferon beta bioavailability muscle albumin
     ; proteinase inhibitor interferon beta bioavailability
ΙT
     Muscle, metabolism
        (interferon-\beta of humans inactivation by,
        albumin and proteinase inhibitors effect on)
     Albumins, biological studies
ΙT
     RL: BIOL (Biological study)
        (muscle inactivation of human interferon-β
        inhibition by)
ΙT
     Interferons
     RL: BIOL (Biological study)
        (β, muscle inactivation of human, albumin and
        proteinase inhibitors effect on)
ΙT
     138674-34-7, Cysteine proteinase inhibitor 139691-92-2, Serine
     proteinase inhibitor
     RL: BIOL (Biological study)
        (muscle inactivation of human interferon-\beta
```

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المتحادث

-7.

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L66 ANSWER 21 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
ΑN
     1991:478932 HCAPLUS
DN
     115:78932
ΕĎ
     Entered STN: 23 Aug 1991
     Stable formulations of lipophilic recombinant proteins
ΤI
IN
     Fernandes, Peter M.; Taforo, Terrance
PA
     Cetus Corp., USA
     U.S., 20 pp. Cont.-in-part of U.S. Ser. No. 752,403.
SO
DT
     Patent
LΑ
     English
     ICM A61K037-02
IC
     ICS A61K045-02
NCL
     424085200
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 16
FAN.CNT 3
                                           APPLICATION NO.
                                                             DATE
    PATENT NO.
                      KIND DATE
                                                            -----
                                           ______
                           _____
                            19910212
                                           US 1985-775751
                                                            19850913 <--
     US 4992271
                      Α
                            19840731
                                           US 1983-495896
                                                            19830518 <--
     US 4462940
                      Α
     CA 1339707
                      A1 19980310
                                           CA 1986-516417
                                                            19860820 <--
                      A1
                          19870319
                                           AU 1986-62642
                                                            19860912 <--
     AU 8662642
                      В2
                            19891123
     AU 590896
                                          EP 1986-307070
     EP 215658
                      A2
                            19870325
                                                            19860912 <--
     EP 215658
                      A3
                            19890208
     EP 215658
                       В1
                            19940601
         R: AT, BE, CH, DE, FR, GB, IT, LI, NL, SE
     AT 106247
                      E
                            19940615
                                           AT 1986-307070
                                                             19860912 <--
     JP 62067032
                       A2
                            19870326
                                           JP 1986-215063
                                                             19860913 <--
     JP 06004542
                       В4
                            19940119
     US 5643566
                       Α
                            19970701
                                           US 1995-474769
                                                             19950607 <--
PRAI US 1982-422421
                            19820923
                                      <--
     US 1983-495896
                            19830518
                                     <--
     US 1984-592077
                            19840323
                                      <--
     US 1985-752403
                            19850705. <--
     US 1985-775751
                            19850913
                                      <--
     EP. 1986-307070
                            19860912
                                      <--
     US 1986-923425
                            19861027
                                      <--
     US 1992-865411
                            19920507
                                      <--
     US 1994-266832
                            19940628
                                      <--
     An improved process for recovering and purifying lipophilic
AB
     {\tt recombinant} proteins such as human {\tt \beta} -
     interferon and interleukin-2 (IL-2) from their hosts yields a
     protein preparation which is formulated into a stable pharmaceutical
composition
     having a therapeutically effective amount of the biol. active
     recombinant lipophilic protein dissolved in a nontoxic, inert,
     therapeutically compatible aqueous based carrier medium at a pH of 6.8 to 7.8.
     The medium also contains a stabilizer for the protein, such as human serum
     albumin and human plasma protein fraction. IL-2 produced by
     recombinant Escherichia coli was purified by a series of steps and
     formulated with human serum albumin (final concentration 2.5%) at pH
ST
     interleukin Escherichia albumin stabilizer; interferon
     recombinant albumin formulation
ΙT
     Escherichia coli
        (beta-interferons and interleukin 2 from)
ΙT
     Proteins, biological studies
     RL: BIOL (Biological study)
        (of blood plasma, as stabilizers for recombinant interleukin
        2-containing pharmaceutical compns.)
```

```
ΙT
     Pharmaceutical dosage forms
        (of recombinant \beta -interferons and
        interleukin 2, stabilizers in, albumins and sugars as)
     Albumins, biological studies
ΤТ
     RL: BIOL (Biological study)
        (stabilizers, for recombinant interleukin 2-containing
        pharmaceutical compns.)
     Lymphokines and Cytokines
ΙT
     RL: BIOL (Biological study)
        (interleukin 2, recombinant, from Escherichia coli,
        stabilized formulations of, albumins and sugars in)
ΙT
     Interferons
     RL: BIOL (Biological study)
        (β , recombinant, from Escherichia coli,
        stabilized formulations of, albumins and sugars in)
ΙT
     69-65-8, Mannitol
     RL: BIOL (Biological study)
        (stabilizer, for recombinant interleukin-2 containing
        pharmaceutical composition)
     50-99-7, Dextrose, biological studies
ΙT
     RL: BIOL (Biological study)
        (stabilizer, for recombinant \beta -
        interferon-containing pharmaceutical composition)
L66 ANSWER 22 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
     1990:153049 HCAPLUS
AN
DN
     112:153049
ΕD
     Entered STN: 28 Apr 1990
ΤI
     Use of human serum albumin signal peptide in recombinant
     protein manufacture and secretion with yeast
     Hayasuke, Naofumi; Nakagawa, Yukimitsu; Ishida, Yutaka; Okabayashi, Ken;
ΙN
     Murakami, Kohji; Tsutsui, Kiyoshi; Ikegaya, Kazuo; Minamino, Hitoshi;
     Ueda, Sadao; et al.
PΑ
     Green Cross Corp., Japan
SO
     Eur. Pat. Appl., 35 pp.
     CODEN: EPXXDW
DΤ
     Patent
     English
LA
IC
     ICM C12N015-00
     ICS C12P021-00
CC
     3-4 (Biochemical Genetics)
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
     ______
                      ____
                           -----
                                           ______
                                                            _____
                                                            19880503 <--
PT
     EP 319641
                      A1
                            19890614
                                           EP 1988-107087
                            19930922
     EP 319641
                       В1
         R: BE, CH, DE, ES, FR, GB, IT, LI, NL, SE
     JP 02167095
                      A2
                            19900627
                                           JP 1988-103339
                                                            19880426 <--
                            19980827
     JP 2791418
                       В2
     CA 1326217
                       Α1
                            19940118
                                           CA 1988-565766
                                                            19880503 <---
                                           ES 1988-107087
                       Т3
                            19941116
                                                            19880503 <--
     ES 2059428
                       В1
                            19970414
                                           KR 1988-5553
                                                            19880513 <--
     KR 9705250
                                                            19950522 <--
     US 5503993
                       Α
                            19960402
                                           US 1995-445783
PRAI JP 1987-306674
                       Α
                            19871202
                                      <--
     JP 1988-45605
                            19880226
                                      <--
                       Α
     US 1988-190553
                       В1
                            19880505
                                      <--
                       В1
                            19920630
                                      <--
     US 1992-913785
OS
     MARPAT 112:153049
     A method for producing and secreting proteins with yeast comprises
AB
     transformation of the yeast with a chimeric gene for a human
    'albumin signal peptide and the coding sequence for the desired
     protein and expression of the gene. Plasmid pNH008, containing the GAL1
```

promoter linked to a synthetic human serum albumin signal

والتترز

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المتأثثة أستري

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sequence fused to the mature human serum albumin gene
     and the pho5 terminator, was constructed. Saccharomyces cerevisiae AH22
     transformed with this plasmid produced 160 mg {\tt albumin}/{\tt L} culture
     medium after 48 h incubation.
     protein secretion yeast albumin signal peptide; Saccharomyces
ST
    human albumin manuf secretion
     Saccharomyces cerevisiae
ΙT
        (human serum albumin manufacture and secretion with,
        albumin signal peptide in)
     Molecular cloning
IT
        (in yeast, human serum albumin signal sequence in)
     Albumins, preparation
IT
     RL: PREP (Preparation)
        (manufacture of, of human, with yeast, human serum albumin signal
        peptide in)
IT
     Lymphokines and Cytokines
     RL: PROC (Process)
        (manufacture of, with yeast, human serum albumin signal peptide
        in)
IT
     Protein sequences
        (of albumin signal peptide analogs, of human)
ΙT
        (recombinant protein secretion from, signal peptide of human
        serum albumin in)
     Deoxyribonucleic acid sequences
ΙT
        (albumin-specifying, signal peptide analog, of human)
IT
     Gene and Genetic element
     RL: BIOL (Biological study)
        (chimeric, for signal sequence of human serum albumin
        and desired protein, expression in yeast of, protein secretion in
        relation to)
     Plasmid and Episome
ΙT
        (pNH008, chimeric human serum albumin signal
        peptide-albumin gene on, expression in Saccharomyces
        cerevisiae of, albumin secretion in relation to)
     Peptides, biological studies
ΙT
     RL: BIOL (Biological study)
        (signal, of human serum albumin, protein secretion from
        recombinant yeast using)
ΙT
     Gene and Genetic element, animal
        (signal sequence, of human serum albumin gene, protein
        secretion from yeast in relation to)
     Interferons
TΤ
     RL: PROC (Process)
        (\alpha , manufacture of, with yeast, human serum \textbf{albumin}
        signal peptide in)
IT
     Interferons
     RL: PROC (Process)
        (\beta , manufacture of, with yeast, human serum albumin
        signal peptide in)
TΤ
     Interferons
     RL: PROC (Process)
        (γ, manufacture of, with yeast, human serum albumin signal
        peptide in)
                    125677-91-0P 125677-92-1P
                                                   125677-93-2P
                                                                   125677-94-3P
IT
     125677-90-9P
     125677-95-4P
     RL: PREP (Preparation)
        (human serum albumin signal peptide derivative,
        recombinant protein manufacture and secretion with yeast in relation
        to)
     125677-89-6P
ΙT
     RL: PREP (Preparation)
```

(human serum albumin signal peptide, recombinant

```
protein manufacture and secretion with yeast in relation to)
     9001-27-8P, Factor VIII 9002-72-6P, Growth hormone
IT
                                  9039-53-6P, Urokinase
                                                           11096-26-7P,
     Insulin, biological studies
     Ervthropoietin
                     62683-29-8P, Colony-stimulating factor
                                                             85637-73-6P,
     Atriopeptin
     RL: IMF (Industrial manufacture); PREP (Preparation)
        (manufacture and secretion of, with yeast, human serum albumin
        signal peptide in relation to)
ΙT
     126115-99-9P
     RL: PREP (Preparation)
        (nucleotide sequence encoding human serum albumin signal
        peptide, recombinant protein manufacture and secretion with yeast
        in relation to)
    ANSWER 23 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
ΑN
     1989:639534 HCAPLUS
DN
     111:239534
ED
     Entered STN: 23 Dec 1989
     Pharmaceutical compositions containing recombinant
TΙ
     interferon-B
     Taforo, Terrance; Thomson, Jody; Shaked, Ze'ev; Hershenson, Susan;
ΙN
     Thomson, James W.; Stewart, Tracy
PΑ
     Cetus Corp., USA
     PCT Int. Appl., 80 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM A61K047-00
     ICS A61K045-02
     63-6 (Pharmaceuticals)
FAN.CNT 2
     PATENT NO.
                      KIND DATE
                                          APPLICATION NO. DATE
                                           _____
PΙ
     WO 8902750
                     A1
                            19890406
                                           WO 1988-US3313
                                                            19880926 <--
         W: AU, DK, JP, NO
         RW: AT, BE, CH, DE, FR, GB, IT, LU, NL, SE
                            19930202
                                           US 1987-100679
                                                            19870929 <--
     US 5183746
                      Α
     AU 8825351
                            19890418
                                           AU 1988-25351
                                                            19880926 <--
                      Α1
PRAI US 1987-100679
                            19870929 <--
     US 1986-923423
                            19861027
                                     <--
     WO 1988-US3313
                            19880926 <--
     A stable parenteral composition in liquid or lyophilized form comprises a
AB
     recombinant interferon-\beta (IFN-.
     beta.) protein dissolved in an inert carrier medium containing
     nonionic polymeric surfactants as a solubilizer/stabilizer. The
     surfactants include polyoxyethylene sorbitan fatty acid esters, a mixture of
     ethoxylated fatty alc. ethers and lauryl ether, ethoxylated octylphenol, a
     mixture of ethoxylated or propoxylated alcs., polyethylene glycol
     monooleate, ethoxylated phenol, and propylene oxide-ethylene oxide block
     copolymers. The composition further comprises addnl. bulking/stabilizing
     agents, such as dextrose. An IFN-\beta analog
     designated as \text{IFN-}\beta ser17 was recovered from
     Escherichia coli culture media and stabilized by adding 0.15% Trycol
     LAL-12 and pH was adjusted to 7.0 with NaOH. A bulking/stabilizing agent,
     i.e., 5% dextrose, was then added and the solution was sterile-filtered,
     aseptically filled into vials, and lyophilized. The IFN-.
     beta. formulations of this invention contain very low levels of
     aggregates and other potentially immunogenic characterisitcs and minimal
     or no strong solubilizing agents, such as SDS, and they are nontoxic and
     have good shelf life.
ST
     interferon beta surfactant solubilizer injection;
     lyophilization interferon beta stability
     Solubilizers
ΙT
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Stabilizing agents
        (nonionic surfactants and sugars as, for interferon
        β -containing parenteral compns.)
ΙT
    Albumins, biological studies
    RL: BIOL (Biological study)
        (parenteral interferon-β composition containing
        nonionic surfactants and, as stabilizer)
ΙT
    Carbohydrates and Sugars, biological studies
    RL: BIOL (Biological study)
        (parenteral interferon-β composition containing
       nonionic surfactants and, as stabilizers)
ΙT
    Surfactants
        (nonionic, parenteral interferon-β composition
        containing, as stabilizers)
     Pharmaceutical dosage forms
IT
        (parenterals, containing \beta -interferons, nonionic
        surfactants and sugars in, as solubilizers/stabilizers)
IT
     Interferons
    RL: BIOL (Biological study)
        (β , parenteral compns. containing, solubilizers/stabilizers
        for, nonionic surfactants and sugars as)
     50-70-4, Sorbitol, biological studies 50-99-7, Dextrose, biological
ΙT
               56-81-5, Glycerol, biological studies 69-65-8, Mannitol
    87-89-8, Inositol 151-21-3, Sodium dodecyl sulfate, biological studies
    RL: BIOL (Biological study)
        (parenteral interferon-\beta composition containing
        nonionic surfactants and, as stabilizer)
     9002-92-0, Ethoxylated lauryl alcohol 9002-93-1, Triton X305
IT
     9004-78-8, Ethoxylated phenol 9004-96-0 9005-64-5, Polyoxyethylene
     sorbitan monolaurate 9005-65-6 9036-19-5, Ethoxylated octylphenol
     12616-49-8, Plurafac C17
                               106392-12-5, Propylene oxide-ethylene oxide
    blocker copolymer
     RL: BIOL (Biological study)
        (parenteral interferon-\beta composition containing, as
        stabilizer)
L66 ANSWER 24 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
ΑN
    1989:18548 HCAPLUS
DN
    110:18548
ΕD
    Entered STN: 21 Jan 1989
    Method for treatment of essential (hemorrhagic) thrombocythemia with human 3
TI
    Delwiche, Francis; Flament-Grivegnee, Jocelyn; Gangji, Diamond; Monsieur,
ΙN
    Rita; Stryckmans, Pierre; Velu, Thierry; Wybran, Joseph
    Boehringer Ingelheim International G.m.b.H., Fed. Rep. Ger.
PΑ
SO
    U.S., 4 pp.
    CODEN: USXXAM
DT
    Patent
LA
    English
    ICM A61K045-02
IC
NCL
    424085000
    1-8 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 1
    PATENT NO.
                     KIND DATE
                                          APPLICATION NO.
                                                            DATE
                     ---- ------ ,
     ______
                                        ------
    US 4743445
                    A 19880510
                                           US 1985-758729 19850725 <--
ΡI
PRAI US 1985-758729
                          19850725 <--
    Essential thrombocythemia is treated by administration of an effective
     amount of human \alpha -interferon. Patients with
     essential thrombocythemia were given i.m. injections of 5 + 106 IU
     recombinant human interferon-\alpha 2 (Arg)
     (I)/day for 30 days. After 15 days, the dose was doubled if the results
```

of the treatment were insufficient. After 30 days, the same dose was given twice a week as a maintenance dose. In all patients the number of thrombocytes returned to normal. A parenteral formulation comprises I 5 + 106 IU, isotonic phosphate buffer (pH 7) q.s., human serum albumin 20.0 mg, and water for injection 1.0 mL. STessential thrombocythemia alpha interferon ΙT Blood platelet (α -interferon of human effect on) ΙT Blood platelet (disease, essential thrombocythemia, treatment of, with  $\alpha$ -interferon of human) Interferons RL: BIOL (Biological study)  $(\alpha$  , essential thrombocythemia treatment with, of human) . 118104-04-4 RL: BIOL (Biological study) (essential thrombocythemia treatment with) ANSWER 25 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN L66 ΑN **1988:562850** HCAPLUS DN 109:162850 ΕD Entered STN: 12 Nov 1988 TΙ Recombinant human interferon alpha-2a: delivery to lymphoid tissue by selected modes of application ΑU Supersaxo, Andreas; Hein, Wayne; Gallati, Harald; Steffen, Hans CS Preclin. Dev., F. Hoffmann-La Roche und Co. Ltd., Basel, Switz. SO Pharmaceutical Research (1988), 5(8), 472-6 CODEN: PHREEB; ISSN: 0724-8741 DT Journal LA English CC 1-2 (Pharmacology) Following s.c. or injection device (i.d.) administration, AΒ recombinant human interferon  $\alpha$  -2a (rIFN  $\alpha$ -2a) of mol. weight 19,000 was absorbed mainly by the lymphatics. This results in high rIFN  $\alpha$ -2a levels in the lymphoid tissue which drains the application site, while blood plasma levels are relatively low. The maximum measured concns. of rIFN  $\alpha$ -2a in the efferent popliteal lymph varied by a factor of 105 between intradermal/s.c. and i.v. administration and was affected neither by the infusion rate nor by the coadministration of albumin. This may help to improve the mode of administration and therapeutic efficacy of protein drugs whose targets are lymphoid cells. interferon  $\alpha$  2a delivery lymph gland ST ΙT Lymphatic system (interferon  $\alpha$  -2a absorption by, after parenteral administrations) ΙT Albumins, biological studies RL: BIOL (Biological study) (interferon  $\alpha$  -2a delivery to lymphoid tissue in relation to) ΙT Lymph gland (interferon  $\alpha$  -2a delivery to, parenteral administration routes for) ΙT Interferons RL: BIOL (Biological study)  $(\alpha - 2a, delivery to lymphoid tissue of$ recombinant, parenteral administration routes for) ANSWER 26 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN L66 1987:583557 HCAPLUS ANDN 107:183557 Entered STN: 14 Nov 1987 ΕD

ور تراتيخ

والمتقال

ΤI

Improved formulation for recombinant  $\beta$  -

```
interferon with protein or sugar stabilizer
     Hanisch, Wolfgang Helmut; Taforo, Terrance; Fernandes, Peter Michael
ΙN
PΑ
     Cetus Corp., USA
     Eur. Pat. Appl., 34 pp.
SO
     CODEN: EPXXDW
DT
     Patent
LA
     English
     ICM A61K045-02
IC
     ICS A61K047-00; C07K003-02; C12P021-02
     63-6 (Pharmaceuticals)
CC
     Section cross-reference(s): 3
FAN.CNT 3
     PATENT NO.
                       KIND DATE
                                             APPLICATION NO. DATE
                                             _____
                                             EP 1986-307070
                                                               19860912 <--
                       A2
                             19870325
     EP 215658
PΙ
     EP 215658
                             19890208
                       Α3
     EP 215658
                       В1
                            19940601
         R: AT, BE, CH, DE, FR, GB, IT, LI, NL, SE
     US 4992271
                                             US 1985-775751
                                                               19850913 <--
                      Α
                            19910212
     AT 106247
                                                              19860912 <--
                             19940615
                                             AT 1986-307070
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PRAI US 1985-775751
                             19850913
                                       <--
     US 1982-422421
                             19820923
                                       <--
     US 1983-495896
                             19830518
                                       <--
     US 1984-592077
                             19840323
                                        <--
     US 1985-752403
                             19850705
                                        <--
     EP 1986-307070
                             19860912
     Recombinant \beta-human interferon (.beta
AΒ
     .-HIFN) is dissolved in a non-toxic, inert, therapeutically compatible aqueous
     carrier, at a pH of 2-4. The solution contains a stabilizer for the
     \beta\textsc{-HIFN},\ \textsc{particularly}\ \textsc{human}\ \textsc{plasma}\ \textsc{protein}\ \textsc{fraction},\ \textsc{human}\ \textsc{serum}
     albumin, or mannitol. This formulation results in very low sodium
     dodecyl sulfate levels. \beta -Interferon 0.25 mg/mL
     was formulated using 2.5% plasma protein fraction at pH 3-4, incubated
     15-45 min.; the pH was adjusted to 7.3-7.5. At this pH, the solns. were
     very clear. The use of 5.0% human serum albumin also gave clear
     solns., whereas 2.5% HSA resulted in slightly hazy solns.
     interferon formulation protein solubilization; stabilizer
ST
     recombinant beta interferon
     Albumins, biological studies
ΙT
     RL: BIOL (Biological study)
         (human, stabilizer for recombinant \beta-human
        interferon)
     Proteins, specific or class, biological studies
TΨ
     RL: BIOL (Biological study)
         (of blood plasma, as stabilizer for recombinant \beta-human
        interferon)
ΙT
     Recombination, genetic
         (of \beta -interferon, purification and formulation for)
IT
     Interferons
         (\beta -, recombinant, stabilization of, in
        formulation)
     151-21-3, Sodium dodecyl sulfate, biological studies
IT
     RL: PRP (Properties)
         (reduced levels of, in formulations of \beta -
        interferon)
     50-99-7, Dextrose, biological studies
                                               69-65-8, Mannitol
TΤ
     RL: BIOL (Biological study)
         (stabilizer, for recombinant \beta -
        interferon-containing pharmaceutical composition)
L66 ANSWER 27 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
     1987:464710 HCAPLUS
AN
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107:64710

DN

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Entered STN: 21 Aug 1987
ED
ΤI
     Potency stability of recombinant (serine-17) human
     interferon-β
     Geigert, John; Ziegler, Diana L.; Panschar, Barbara M.; Creasey, Abla A.;
AU
     Vitt, Charles R.
CS
     Dep. Tech. Dev., Cetus Corp., Emeryville, CA, 94608, USA
SO
     Journal of Interferon Research (1987), 7(2), 203-11
     CODEN: JIREDJ; ISSN: 0197-8357
DT
     Journal
     English
LA
CC
     63-3 (Pharmaceuticals)
     The antiviral activity of Escherichia coli-derived (serine-17) human
AΒ
     interferon-\beta , formulated with human serum
     albumin, is stable for 2 yr when lyophilized and stored under
     refrigeration. This product shows an Arrhenius line fit for the stability
     of its activity when tested at multiple isothermal temps. (25-80°).
     In both isothermal and non-isothermal elevated temperature studies, increasing
     the level of human serum albumin in the formulation results in
     increased thermal stability.
     interferon serine 17 recombinant formulation stability
ST
     Kinetics of decomposition
        (of recombinant human β -interferon
        in albumin formulation)
     Albumins, uses and miscellaneous
ΙT
     RL: USES (Uses)
        (β -interferon recombinant serine-17
        stabilization by formulation with human)
ΙT
     Interferons
        (\beta -, stability of recombinant serine-17, in
        human serum albumin formulation)
    ANSWER 28 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN
L66
     1986:174635 HCAPLUS
AN
DN
     104:174635
     Entered STN: 17 May 1986
ED
     Interferon solubilization with amino acids
TI
     Kato, Yasuki; Hayakawa, Eiji; Furuya, Kunitoshi; Kondo, Akira
ΙN
     Kyowa Hakko Kogyo Co., Ltd., Japan
PΑ
SO
     Eur. Pat. Appl., 14 pp.
     CODEN: EPXXDW
DT
     Patent
     English
LA
     ICM A61K045-02
IC
     63-3 (Pharmaceuticals)
     Section cross-reference(s): 15
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO. DATE
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                                           -----
                                                           _____
     EP 163111
                            19851204
                                           EP 1985-104849
                                                            19850422 <--
                      A2
     EP 163111
                     А3
                            19870930
                            19901003
     EP 163111
                     В1
        R: DE, FR, GB, IT
     JP 60243028
                     A2
                            19851203
                                           JP 1984-86972
                                                            19840428 <--
     JP 05058000
                       B4
                            19930825
     CA 1264665
                            19900123
                                           CA 1985-479841
                                                            19850423 <--
                       Α1
     US 4675183
                      Α
                            19870623
                                           US 1985-726971
                                                            19850425 <--
PRAI JP 1984-86972
                            19840428
                                     <--
     Interferon is solubilized by addition of 5 + 10-6 - 5 +
     10-3 mol amino acid/106 units interferon. The amino acid may be
     arginine, histidine, lysine, hydroxylysine, ornithine, glutamine,
```

 $\gamma$ -aminobutyric acid,  $\epsilon$ -aminocaproic acid, or a salt of these

compds. Thus, 5 mg serum albumin, 5 mg NaCl, 30 mg arginine-HCl, and 3 + 106 units of  $\gamma$ - interferon were

- F. .

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LA CC

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(albumin effect on recombinant  $\alpha 2$ -

mixed with 2 mL H2O, and freeze-dried. The product was dissolved in 5 mL  $\,$ H2O, held 6 h at 25°, and the absorbance was measured at 400 nm. The amount of  $\gamma$ - interferon that remained in solution was 98%. This solubilization may be used to facilitate the isolation and purification of interferon produced by recombinant DNA technol. interferon solubilizer amino acid; arginine interferon solubilization Solubilizers (amino acids, for interferon) Amino acids, uses and miscellaneous RL: PRP (Properties) (interferons solubilization by) Interferons  $(\alpha -, solubilization of, with amino acids)$ Interferons  $(\beta$  -, solubilization of, with amino acids) Interferons  $(\gamma$ -, solubilization of, with amino acids) 56-87-1, properties 60-32-2 70-26-8 56-85-9, properties 74-79-3, properties properties 657-27-2 1119-34-2 1190-94-9 60259-81-6 2835-81-6 RL: PRP (Properties) (interferons solubilization by) ANSWER 29 OF 29 HCAPLUS COPYRIGHT 2004 ACS on STN L66 1986:86802 HCAPLUS 104:86802 Entered STN: 22 Mar 1986 The lymphatic route - II. Pharmacokinetics of human recombinant interferon- $\alpha$  2 injected with albumin as a retarder in rabbits Bocci, Velio; Muscettola, Michela; Naldini, Antonella; Bianchi, Enrica; Segre, Giorgio Inst. Gen. Physiol., Univ. Siena, Siena, 53100, Italy General Pharmacology (1986), 17(1), 93-6 CODEN: GEPHDP; ISSN: 0306-3623 Journal English 15-5 (Immunochemistry) An investigation was conducted to define whether multisite s.c. AB . administration in unanesthetized, unrestrained rabbits of human recombinant interferon- $\alpha$  2 (rec. IFN- $\alpha$  2) either in saline, human albumin (ALB) solution (4, 7, and 10% final concns.), or in a solution containing 75 units of hyaluronidase, modified the pharmacokinetic parameters calculated from the IFN plasma level. Plasma disappearance rates of rec. IFN-. alpha.2 were measured in rabbits after i.v. administration and the kinetics was adequately represented by a 3-compartment mammillary model. This model was the basis for evaluating the absorption and distribution of rec. IFN- $\alpha$  2 after s.c. administration. The increase of ALB concentration (from 4 to 10%) caused a significant reduction of the plasma IFN maximum clearance, while both the mean residence time and the release time of IFN increased linearly with the ALB concentration The data support the postulation that s.c. administration of albumin acts as an interstitial fluid expander and may favor absorption of IFN via lymphatics rather than blood capillaries. Improvement of therapeutic index of IFN by using this route remains to be shown in clin. trials. interferon alpha pharmacokinetics albumin Lymphatic system

interferon pharmacokinetics in relation to, of humans and laboratory

animals)

IT Blood plasma

( $\alpha$ 2- interferon pharmacokinetics in, albumin effect on, in humans and laboratory animals)

IT Albumins

RL: BIOL (Biological study)

( $\alpha 2$ - interferon pharmacokinetics response to, of humans and laboratory animals)

IT Interferons

RL: BIOL (Biological study)

( $\alpha$  2-, pharmacokinetics of recombinant

, albumin effect on, of humans and laboratory animals)

=> => fil wpix

FILE 'WPIX' ENTERED AT 16:25:05 ON 02 FEB 2004

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FILE LAST UPDATED:
MOST RECENT DERWENT UPDATE:

· 28 JAN 2004

<20040128/UP>

200407

<200407/DW>

`<<<

DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> NEW WEEKLY SDI FREQUENCY AVAILABLE --> see NEWS <<<

- >>> SLART (Simultaneous Left and Right Truncation) is now
   available in the /ABEX field. An additional search field
   /BIX is also provided which comprises both /BI and /ABEX <<</pre>
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<<
- >>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

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- >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://thomsonderwent.com/coverage/latestupdates/ <<<
- >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER GUIDES, PLEASE VISIT: http://thomsonderwent.com/support/userguides/
- >>> ADDITIONAL POLYMER INDEXING CODES WILL BE IMPLEMENTED FROM DERWENT UPDATE 200403.

  THE TIME RANGE CODE WILL ALSO CHANGE FROM 018 TO 2004.

  SDIS USING THE TIME RANGE CODE WILL NEED TO BE UPDATED.

  FOR FURTHER DETAILS: http://thomsonderwent.com/chem/polymers/ <<<
- => d all abeq tech abex tot

L88 ANSWER 1 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN

AN 2003-421048 [39] WPIX

DNC C2003-110745

TI New hybrid polypeptide, useful for sequestering and/or purifying a polypeptide of interest.

DC B04 D16

IN THOMAS, T; TILLETT, D

PA (PROT-N) PROTIGENE PTY LTD

CYC 101

- -

PI WO 2003018616 A1 20030306 (200339)\* EN 66p C07K001-14

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR IE IT KE LS LU

MC MW MZ NL OA PT SD SE SK SL SZ TR TZ UG ZM ZW

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ADT WO 2003018616 A1 WO 2002-AU1159 20020827

PRAI AU 2001-7298 20010827

ICM C07K001-14

C07K001-36; C07K019-00; C12N009-00; C12N015-63

WO2003018616 A UPAB: 20030619 AΒ

NOVELTY - A hybrid polypeptide comprises a polypeptide of interest linked to a polymerizable polypeptide, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) sequestering and/or purifying a polypeptide of interest;
- (2) a hybrid nucleic acid comprising a nucleic acid encoding the hybrid polypeptide;
- (3) a library comprising several hybrid nucleic acids, polypeptides or vectors;
  - (4) a vector comprising the hybrid nucleic acid;
- (5) a cell transformed or transfected with the hybrid nucleic acid or vector; and
  - (6) purifying a polypeptide of interest.

USE - The hybrid polypeptide is useful for sequestering and/or purifying a polypeptide of interest (claimed). Dwq.0/9

CPI FS

AB; DCN FΑ MC CPI: B04-B04C; B04-C01; B04-E08; B04-F0100E; B04-G01; B04-H01; B04-H02B; B04-H04; B04-H05; B04-H19; B04-J01; B04-J02; B04-J05; B04-J10; B04-L04; B04-L05; B04-L06; B04-L07; B04-N03; B04-N04; B04-N06; B04-N08; B11-B; D05-C11; D05-H12A; D05-H12E; D05-H13; D05-H14; D05-H17C UPTX: 20030619

TECH

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م والمراجعة

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Polypeptide: The hybrid polypeptide is produced in vivo. It is linked to a support, comprising the polymerizable polypeptide. The support polymerizable polypeptide comprises a polymerizable polypeptide identical to the hybrid polypeptide, or its variant. The polypeptide of interest is linked to the polymerizable polypeptide by fusing the polypeptide of interest directly to the polymerizable polypeptide or by a linker polypeptide. It is prokaryotic or eukaryotic in origin. It is a synthetic polypeptide. It comprises endonuclease, a methylase, an oxidoreductase, a transferase, a hydrolase, a lysase, an isomerase, a ligase, a storage polypeptide, a fertitin, an ovalbumin, a transport protein, hemoglobin, serum albumin or ceruloplasmin, an antigen, an antigenic determinant for use in the preparation of vaccines or diagnostic agents, a protective protein, a defense protein, thrombin, fibrinogen, binding proteins, antibodies, immunoglobulins, a human growth hormone, somatostatin, prolactin, estrange, progesterone, melanocyte, thyrotropin, calcitonin, gonadotropin, insulin, a hormone identified as being involved in the immune system, interleukin 1, interleukin 2, colony simulating factor, macrophage-activating factor, interferon, a structur al element, collagen, elastin, alpha-keratin, glyco-protein, virus-protein and muca-protein. The linker polypeptide comprises a recognition site for a proteolytic agent and a multiple cloning site. It also comprises a spacer polypeptide of sufficient length to allow or enhance cleavage of the polypeptide of interest from the polymerizable polypeptide, or to avoid unfavorable steric interference between the polypeptide of interest and the polymerizable polypeptide.

The recognition site comprises an amino acid sequence consisting of:

- (a) Leu-Glu-VaI-Leu-Phe-Gln-Gly-Pro;
- (b) Leu-Val-Pro-Arg-Gly-Ser;

(c) Ile-Glu-Gly-Arg; or

1

(d) Asp-Asp-Asp-Lys.

The chemical capable of proteolytic activity is cyanogen bromide. The polypeptides are linked by antibody interaction, which is achieved by:

- (a) attaching an antibody specific for the polypeptide of interest to the polymerizable polypeptide; or
- (b) using a bi-specific antibody directed to both the polypeptide of interest and the polymerizable polypeptide.

The polymerizable polypeptide is a polypeptide that naturally polymerizes with itself. It is tubulin or actin. It is an FtsZ or Escherichia coli FtsZ protein or its variant. The variant Escherichia coli FtsZ protein comprises replacement of the aspartate residue at position 212 of the protein with a cysteine or asparagine residue. The variant FtsZ protein comprises a mutation selected from replacement of alanine by threonine at position 70, replacement of aspartate by alanine at position 209 or replacement of aspartate by alanine at position 269. The polymerizable polypeptide requires an intermediary polypeptide or other molecule in order to polymerize.

Preferred Method: Sequestering and/or purifying a polypeptide of interest comprises polymerizing the hybrid polypeptide under controlled chemical and/or physical conditions. It is polymerized by a change in temperature and by the addition of an agent that induces polymerization. The polymerization inducing agent is GTP, ATP and/or a cation. The cation comprises magnesium, calcium, nickel, cobalt, zinc or manganese. The polymerized hybrid polypeptide is purified by a first purification step, which may be the only purification step or may be followed by further purification steps. The first purification step purifies the polymerized hybrid polypeptide by physical techniques discriminating on the basis of size and/or weight. The polymerized hybrid polypeptide is also purified by centrifugation, differential sedimentation, filtration, dialysis and/or flow sorting, where the polymerized hybrid polypeptide is isolated. After the first purification step the polymerized hybrid polypeptide is dissociated. The dissociation is achieved by removal of the agent which induces polymerization and/or incubation of the polymerized hybrid polypeptide at a suitable temperature. The dissociated hybrid polypeptide is purified by a second purification step, which comprises purification of the hybrid polypeptide on the basis of size and/or weight. The polymerization, dissociation and purification of the polymerizable hybrid polypeptide are repeated so that substances larger and smaller than the hybrid polypeptide are removed. The polymerizable polypeptide is cleaved from the polypeptide of interest by a proteolytic agent, which does not substantially interfere with the biological or chemical activity of the polypeptide of interest or the polymerizable polypeptide. After the cleavage of the polypeptide of interest from the polymerizable polypeptide, the protease hybrid polypeptide is polymerized. The proteolytic agent comprises 3C-protease from a human rhinovirus type 14 (HRV protease 3C), thrombin, Factor Xa, enterokinase and a chemical capable of proteolytic activity. It is linked to a polymerizable polypeptide to form a protease hybrid polypeptide. The polymerizable polypeptide to which the protease is linked is identical to the polymerizable polypeptide to which the polypeptide of interest is linked, or is a variant of it.

Purifying a polypeptide of interest comprises:

- (a) expressing the hybrid nucleic acid in a cell to produce a hybrid polypeptide comprising the polypeptide of interest and a polymerizable polypeptide;
- (b) polymerizing the hybrid polypeptide;
- (c) purifying the polymerized hybrid polypeptide;
- (d) cleaving the polypeptide of interest from the polymerizable polypeptide; and
- (e) purifying the polypeptide of interest.

ABEX

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UPTX: 20030619

EXAMPLE - No suitable example given.

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ANSWER 2 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN
1.88
AN
     2002-179329 [23]
                        WPIX
CR
     2001-602931 [68]
DNC
     C2002-055553
TΤ
     New albumin fusion proteins with extended shelf life, useful for
     treating leukemia, warts, hepatitis, multiple sclerosis and AIDS,
     comprises therapeutic protein fused to albumin.
     B04 D16
DC
ΙN
     BALLANCE, D J; PRIOR, C P; SADEGHI, H; SLEEP, D; TURNER, A J
PΑ
     (DELZ) DELTA BIOTECHNOLOGY LTD; (PRIN-N) PRINCIPIA PHARM CORP; (BALL-I)
     BALLANCE D J; (PRIO-I) PRIOR C P; (SADE-I) SADEGHI H; (SLEE-I) SLEEP D;
     (TURN-I) TURNER A J
CYC
     96
PΙ
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                                                      C07K014-00
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     AU 2001061024 A 20011030 (200225)
                                                      C07K014-00
     EP 1278767
                   A1 20030129 (200310)
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                                                      C07K014-00
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     US 2003199043 A1 20031023 (200370)
                                                      C12P021-02
     JP 2003530839 W 20031021 (200373)
                                              453p
                                                      C12N015-09
ADT WO 2001079271 A1 WO 2001-US12009 20010412; AU 2001061024 A AU 2001-61024
     20010412; EP 1278767 A1 EP 2001-934875 20010412, WO 2001-US12009 20010412;
     US 2003199043 A1 Provisional US 2000-229358P 20000412, Provisional US
     2000-199384P 20000425, Provisional US 2000-256931P 20001221, US
     2001-832501 20010412; JP 2003530839 W JP 2001-576866 20010412, WO
     2001-US12009 20010412
FDT AU 2001061024 A Based on WO 2001079271; EP 1278767 A1 Based on WO
     2001079271; JP 2003530839 W Based on WO 2001079271
PRAI US 2000-256931P 20001221; US 2000-229358P 20000412; US 2000-199384P
     20000425; US 2001-832501
                                20010412
IC
     ICM C07K014-00; C12N015-09; C12P021-02
          A61K038-00; A61K038-16; A61K038-21; A61K038-43; A61K038-46;
          A61K038-48; A61K038-55; A61K039-395; A61K047-48; A61P001-16;
          A61P015-00; A61P017-12; A61P025-28; A61P031-12; A61P031-14;
          A61P031-18; A61P031-20; A61P035-00; A61P035-02; C07H021-04;
          CO7KO14-52; CO7KO14-56; CO7KO14-745; CO7KO14-75;
          C07K014-76; C07K014-765; C07K014-81; C07K016-00;
          C07K019-00; C12N001-19; C12N001-21; C12N005-06; C12N005-10;
          C12N009-14; C12N009-74; C12N009-99; C12N015-00
     WO 200179271 A UPAB: 20031112
AΒ
     NOVELTY - An albumin fusion protein (I) comprising:
           (a) a therapeutic protein (X) and albumin (A) containing a
     fully defined sequence (S1) of 585 amino acids as given in the
     specification;
           (b) X and a fragment or variants of S1, where the fragment or
     variants has albumin activity; or
          (c) a fragment or variant of X and A, where the fragment or variant
     has a biological activity of X, is new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:
           (1) an albumin fusion protein (II) comprising a peptide
     inserted into A comprising amino acids 54-61, 76-89, 92-100, 170-176,
     247-252, 266-277, 280-288, 362-368, 439-447, 462-475, 478-486 or 560-566
     of S1;
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(2) an **albumin** fusion protein (III) comprising a single chain antibody or its portion and A or its fragment or variant;

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- (3) a composition comprising any of (I)-(III) and a pharmaceutically active carrier;
  - (4) a kit comprising the composition;
- (5) treating a disease or disorder that is modulated by X in a patient comprising administering any of (I)-(III);
- (6) extending the shelf life of X comprising fusing X or its fragment or variant to A or its fragment or variant, sufficient to extend the shelf-life of X compared to the shelf life of X in an unfused state;
- (7) a nucleic acid molecule (IV) comprising a polynucleotide sequence encoding any of (I)-(III);
  - (8) a vector comprising (IV); and
  - (9) a host cell comprising (IV).

ACTIVITY - Cytostatic; dermatological; virucide; anti-HIV; neuroprotective; hepatotropic; antiinflammatory. Tests are described but no results are given in the source material.

MECHANISM OF ACTION - Gene therapy.

USE - The fusion protein is useful for the treatment of hairy cell leukemia, Kaposi's sarcoma, genital warts, anal warts, chronic hepatitis B, chronic non-A, non-B hepatitis, hepatitis C/D, chronic myelogenous leukemia, renal cell carcinoma, bladder carcinoma, ovarian carcinoma, cervical carcinoma, skin cancer, recurrent respirator papillomatosis, non-Hodgkin's lymphoma, cutaneous T-cell lymphoma, melanoma, multiple myeloma, acquired immunodeficiency syndrome (AIDS), multiple sclerosis and glioblastoma. The fusion of albumin extends the shelf life and the in vivo and in vitro biological activity of the therapeutic protein (all claimed).

ADVANTAGE - Therapeutic proteins can be stabilized to extend shelf life and/or retain the protein's activity for extended periods of time in solution, in vivo or in vitro by genetically or chemically fusing the protein to albumin or its fragment or variant. In addition the use of albumin fusion proteins reduces the need to formulate protein solutions with large excesses of carrier proteins to prevent loss of therapeutic protein due to factors such as binding to the container. The extension of shelf life was tested by measuring biological activity (Nb2 cell proliferation) of human albumin-human growth hormone (HA-hGH) fusion protein remaining after incubation in cell culture media for up to 3 weeks at 37 deg. C. At week 3 there was still approx. 95% cell proliferation compared to no activity of unfused hGH (no observed activity by week 2).

Dwg.0/18

FS CPI

MC

TECH

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FA AB; DCN

CPI: **B04-C01G**; B04-E02H; B04-E08; B04-F0100E; B04-G01;

B04-H05A; B04-H19; B04-L05A; B04-N02A; B04-N08;

B14-A02A; B14-A02B1; B14-G01B; B14-H01; B14-N12; B14-N17; B14-S01;

B14-S03A; D05-C12; D05-H12C; D05-H12E; D05-H14; D05-H17C

UPTX: 20020411

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preparation: The fusion proteins can be prepared by standard recombinant techniques. Preferred Fusion Protein: Albumin activity is the ability to prolong the shelf life of X compared to the shelf life of X in an unfused state. Preferably the fragment or variant of (I) comprises amino acids 1-387 of S1. X is chosen from serum cholinesterase, alpha-1 antitrypsin, aprotinin, coagulated complex, von Willebrand factor, fibrinogen, factor VII, factor VIIA activated factor, factor VIII, factor IX, factor X, factor XIII, c1 inactivator, antithrombin III, thrombin, prothrombin, apo-lipoprotein, c-reactive protein, protein C, immunoglobulin and preferably interferon (IFN)-alpha. X or its fragment or variant is fused to the N or C-terminus of A. (I)-(III) comprises a first and second X, where the first X is different from the second X. X is separated from A by a linker. The fusion protein has the formula R1-L-R2, R2-L-R1 or R1-L-R2-L-R1, where:

R1 = X

L = peptide linker; and

R2 = A or its fragment or variant.

The in vitro or in vivo activity of X fused to A is greater than the in vitro or in vivo biological activity of X in an unfused state. The protein is expressed in a glycosylation and protease deficient yeast. Alternatively it is expressed by a mammalian cell in culture. The fusion protein further comprises a secretion leader sequence.

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preparation: The fusion proteins can be produced by standard chemical synthetic techniques.

UPTX: 20020411

ABEX

ADMINISTRATION - 1 microgram/kg/day to 10 mg/kg/day, preferably 0.01-1 mg/kg/day of **albumin** fusion proteins are administered by standard routes.

EXAMPLE - A human albumin-human growth hormone (HA-hGH) fusion protein was prepared. The hGH cDNA was obtained from a human pituitary gland cDNA library by polymerase chain reaction (PCR) amplification. The PCR product was purified and then digested with EcoRl and HindIII. After further purification of the EcoR1-HindIII fragment by gel electrophoresis, the product was cloned into pUC19 digested with EcoR1 and HindIII to give pHGH1. The polylinker sequence of the phagemid pBluescribe (+) (Stratagene) was replaced by inserting an oligonucleotide linker formed by annealing 2 75-mer oligonucleotides between the EcoRl and HindIII sites to form pBST(+). The new polylinker included a unique NotI site. the NotI HA expression cassette of pAYE309 comprising the PRBI promoter, DNA encoding the HA/MFalpha-1 hybrid leader sequence, DNA encoding HA and the ADH1 terminator, was transferred to pBST(+) to form pHA1. The HA sequence was removed from this plasmid by digestion with HindIII followed by religation to form pHA2. Cloning of the hGH cDNA provided the hGH coding region lacking the pro-hGH sequence and the first 8 base pairs (bp) of the mature hGH sequence. In order to construct an expression plasmid for secretion of hGH from yeast, a yeast promoter, signal peptide and the first bp of the hGH sequence were attached to the 5' end of the cloned hGH sequence. The HindIII-SfaNI fragment from pHA1 was attached to the 5' end of the EcoR1/HindIII fragment from pHGHI via 2 synthetic oligonucleotides to generate a double stranded fragment of DNA with sticky ends that can anneal with SfaNI and EcoR1 sticky ends. The HindIII fragment formed was cloned into HindIII digested pHA2 to make pHGH2 such that the hGH cDNA was positioned downstream of the PRBI promoter and HA/MFalpha-1 fusion leader sequence. The NotI expression cassette contained in pHGH2 was cloned into the NotI-digested pSAC35 to make pHGH12. This plasmid comprised the entire 2 micro m plasmid to provide replication functions and the LEU2 gene for selection of transformants. pHGH12 was introduced into S. cerevisiae D88 by transformation and individual transformants were grown for 3 days at 30 degrees C in 10 mL YEPD (1% w/v yeast extract, 2% w/v peptone, 2% w/v dextrose). After centrifugation of the cells, the supernatants were examined by sodium dodecyl sulfate polyacrylamide gel electrophoresis (SDS-PAGE) and were found to contain protein which was of the expected size and recognized by anti-hGHG antiserum on Western blots.

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L88 ANSWER 3 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN
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AN **2001-616754** [71] WPIX

CR 2001-602931 [68]; 2001-611723 [70]; 2001-616755 [71]; 2001-616756 [71]; 2002-010886 [01]; 2003-810996 [76]; 2004-033644 [03]

DNC C2001-184720

Albumin fusion proteins comprising a therapeutic protein and albumin, useful in the treating immune system disorders (e.g. transplant rejection), blood related disorders (e.g. myocardial infarction) and hyperproliferative disorders.

DC B04 D16

IN HASELTINE, W A; ROSEN, C A
PA (HUMA-N) HUMAN GENOME SCI INC

CYC 96

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WO 2001079443 A2 20011025 (200171)* EN 365p
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                   A2 20030115 (200313)
                                        EΝ
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                                             469p
    WO 2001079443 A2 WO 2001-US11924 20010412; AU 2001059063 A AU 2001-59063
ADT
     20010412; EP 1274719 A2 EP 2001-932546 20010412, WO 2001-US11924 20010412;
     JP 2003530846 W JP 2001-577427 20010412, WO 2001-US11924 20010412
    AU 2001059063 A Based on WO 2001079443; EP 1274719 A2 Based on WO
     2001079443; JP 2003530846 W Based on WO 2001079443
PRAI US 2000-256931P 20001221; US 2000-229358P 20000412; US 2000-199384P
     20000425
     ICM C07K001-00; C12N000-00; C12N015-09
IC
         A01N037-18; A61K038-00; A61K038-21; A61K038-28;
          A61K039-395; A61K047-48; A61K048-00; A61P001-16; A61P013-00;
          A61P025-00; A61P031-14; A61P031-18; A61P031-20; A61P035-00;
          A61P035-02; C07K014-47; C07K014-76; C07K019-00;
          C12N001-19; C12N005-10
     WO 200179443 A UPAB: 20040112
AΒ
     NOVELTY - Albumin fusion proteins (P1) comprising a therapeutic
     protein (T1) (or its fragment or variant having the activity of T1) and
    albumin comprising the 585 amino acid sequence (I) defined in the
     specification (or its fragment or variant having albumin
     activity), are new.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:
          (1) a kit comprising a composition containing P1;
          (2) a method of treating a disease or disorder, preferably modulated
     by T1, in a patient, comprising administering P1;
          (3) a method of extending the shelf-life of T1, comprising fusing T1
     or its fragment or variant, to albumin or its fragment or
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- in an unfused state;
  (4) a nucleic acid (N1) comprising a nucleotide sequence encoding P1;
  - (5) a vector comprising N1; and
  - (6) a host cell comprising N1.

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ACTIVITY - Cytostatic; antiinflammatory; antileukemic; antiarthritic; antirheumatic; immunosuppressive; cardiant; nootropic; neuroprotective; antimicrobial; vulnerary.

variant, where the shelf-life of T1 or its fragment or variant as part of a fused protein is extended when compared to T1 or its fragment or variant

To test whether sympathetic neuronal cell viability is supported by an albumin fusion protein, the chicken embryo neuronal survival assay (Senaldi, et al., Proc. Natl. Acad., Sci., U.S.A, 96:11458-63 (1998)). Briefly, motor and sympathetic neurons were isolated from chicken embryos, resuspended in L15 medium (with 10% foetal calf serum (FCS), glucose, sodium selenite, progesterone, conalbumin, putrescine and insulin) and Dulbecco's modified Eagles medium (with 10% FCS, glutamine, penicillin, and 25 mM Hepes buffer (pH 7.2)), respectively and incubated at 37 degrees Centigrade in 5% carbon-dioxide in the presence of different concentrations of the purified fusion protein, as well as negative control lacking any cytokine, After 3 days, neuronal survival was determined by evaluation of cellular morphology, and through the use of the colorimetric assay of Mosmann (Mosmann, T., J. Immunol., Methods, 65:55-63 (1983)). Enhanced neuronal cell viability as compared to the controls lacking cytokine is indicative of the ability of the albumin fusion protein to enhance the survival of neuronal cells.

MECHANISM OF ACTION - Gene therapy.

USE - The albumin fusion proteins are also useful in the treatment, prevention, diagnosis, and/or detection of diseases, disorders such as immune system disorders (e.g. transplant rejection), blood related disorders (e.g. myocardial infarction), hyperproliferative disorders (e.g. childhood acute myeloid leukemia), renal disorders (e.g. qlomerulonephritis), cardiovascular disorders (e.g. arrhythmias), respiratory disorders (e.g. non-allergic rhinitis), neurological diseases (e.g. Alzheimer's disease), endocrine disorders (e.g. pheocytochroma), reproductive system disorders (e.g. syphilis), infectious diseases (e.g. measles), gastrointestinal disorders (e.g. irritable bowel syndrome) and wound healing.

Dwg.0/15

FS CPI

FΑ AB; DCN

CPI: **B04-C01**; B04-E02F; B04-E08; B04-F0100E; B04-F0200E; MCB04-F0900E; B04-F1100E; B04-N02A0E; B14-A01; B14-A02; B14-D01; B14-E10; B14-F01; B14-F02; B14-G01; B14-G02; B14-G03; B14-H01; B14-J01; B14-K01; B14-N10; B14-N17B; B14-S03;

DO5-H12B2; D05-H12E; D05-H14A2; D05-H14B2

UPTX: 20011203

TECH

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TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Fusion Protein: The albumin activity is the ability to prolong the shelf-life of T1 compared to the shelf-life of T1 in an unfused state. The albumin fragment or variant comprises amino acids 1-387 of (I). T1 or its fragment or variant is fused to the C-terminal of the albumin or the C-terminus of the fragment or variant of albumin. Alternatively, T1 or its fragment or variant is fused to the N-terminal of the albumin or the N-terminus of the fragment or variant of albumin. Alternatively, T1 or its fragment or variant is fused to the N-terminus and C-terminus of the albumin , or the N-terminus and C-terminus of the fragment or variant of albumin.

P1 comprises a first T1 or its fragment or variant, and a second T1 or its fragment or variant, where the first T1 is different from the second T1. T1 or its fragment or variant is separated from the albumin or the fragment or variant of albumin by a linker. Preferably, P1 is of the formula (S1), (S2) or (S3). R1-L-R2 (S1);

R2-L-R1 (S2); or

R1-L-R2-L-R1 (S3).

R1 = is T1 or its fragment or variant;

L = is a peptide linker; and

R2 = is albumin comprising the sequence of (I), or its fragment or variant.

The shelf-life of the albumin fusion protein is greater than the shelf-life of T1 or its fragment or variant in an unfused state. The in vitro or in vivo biological activity of T1 or its fragment or variant, fused to albumin or its fragment or variant, is greater than the in vitro or in vivo, respectively, biological activity of T1 or its fragment or variant, in an unfused state.

Alternatively, Pl comprises Tl or its fragment or variant, inserted into an albumin comprising the sequence of (I) or its fragment or variant. Preferably, the albumin comprises residues 54-61, 76-89, 92-100, 170-176, 247-252, 266-277, 280-288, 362-368, 439-447, 462-475, 478-486, or 560-566 of (I). The portion of albumin is sufficient to prolong the shelf-life of T1, or its fragment or variant, as compared to the shelf-life of T1, or its fragment or variant in an unfused state.

The portion of albumin is sufficient to prolong the in vitro and in vivo biological activity of T1 or its fragment or variant, as compared to the in vitro and in vivo biological activity of T1 or its fragment or

ABEX

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variant, in an unfused state. P1 is non-glycosylated and is expressed in yeast which is glycosylation deficient. The yeast may also be protease deficient. Alternatively, Pl is expressed by a mammalian cell in culture. Pl further comprises a secretion leader sequence. UPTX: 20011203 ADMINISTRATION - The albumin fusion proteins can be administered orally, rectally, parenterally, intracisternally, intravaginally, intraperitoneally, topically, bucally, or as an oral or nasal spray. The dosage is 1 microgram/kg/day to 10 mg/kg/day, preferably 0.01 to 1, mg/kd/day. If given continuously, the albumin fusion protein is typically administered at a dose rate of 1-50 micrograms/kg/hour, either by 1-4 injections per day or by continuous subcutaneous infusions. ANSWER 4 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN **2001-611723** [70] WPIX 2001-602931 [68]; 2001-616754 [71]; 2001-616755 [71]; 2001-616756 [71]; 2002-010886 [01]; 2003-810996 [76]; 2004-033644 [03] C2001-182838 New albumin fusion proteins, useful for treating diseases and disorders such as cancer, comprise therapeutic protein fused to albumin. B04 D16 HASELTINE, W A; ROSEN, C A (HUMA-N) HUMAN GENOME SCI INC 96 WO 2001079442 A2 20011025 (200170)\* EN 362p C12N000-00 RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR.TT TZ UA UG US UZ VN YU ZA ZW AU 2001064563 A 20011030 (200219) C12N000-00 A2 20030122 (200315) C12N001-18 EP 1276849 ENR: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR 540p JP 2003531590 W 20031028 (200373) C12N015-09 WO 2001079442 A2 WO 2001-US11850 20010412; AU 2001064563 A AU 2001-64563 20010412; EP 1276849 A2 EP 2001-938994 20010412, WO 2001-US11850 20010412; JP 2003531590 W JP 2001-577426 20010412, WO 2001-US11850 20010412 AU 2001064563 A Based on WO 2001079442; EP 1276849 A2 Based on WO 2001079442; JP 2003531590 W Based on WO 2001079442 PRAI US 2000-256931P 20001221; US 2000-229358P 20000412; US 2000-199384P 20000425 C12N000-00; C12N001-18; C12N015-09 ICM A61K038-00; A61K038-21; A61K039-395; A61K048-00; A61P001-04; A61P001-16; A61P001-18; A61P003-10; A61P005-14; A61P005-40; A61P007-04; A61P007-06; A61P009-00; A61P009-06; A61P009-10; A61P009-12; A61P011-00; A61P011-06; A61P013-00; A61P013-02; A61P013-08; A61P013-12; A61P015-00; A61P015-10; A61P015-18; A61P017-00; A61P017-02; A61P019-00; A61P019-02; A61P019-08; A61P021-00; A61P021-04; A61P025-00; A61P025-08; A61P025-16; A61P025-28; A61P027-02; A61P029-00; A61P031-00; A61P031-12; A61P031-16; A61P031-18; A61P031-22; A61P033-02; A61P033-06; A61P033-12; A61P035-00; A61P035-02; A61P037-00; A61P037-08; A61P039-02; A61P041-00; A61P043-00; C07K014-47; C07K014-76; C07K019-00; C12N001-19; C12N005-10 WO 200179442 A UPAB: 20040112 NOVELTY - An albumin fusion protein (I) comprising a therapeutic protein: X and (a fragment or variant of) albumin comprising a

fully defined sequence (S18) of 585 amino acids as given in the specification, (where the fragment or variant has albumin or

therapeutic protein: X activity) is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) a kit comprising a composition containing (I);
- (2) treating a disease or disorder (that is modulated by therapeutic protein: X or its fragment or variant) comprising administering (I);
- (3) extending the shelf life of therapeutic protein: X comprising fusing therapeutic protein: X or its fragment or variant to albumin or its fragment or variant, sufficient to extend the shelf life of therapeutic protein: X compared to the shelf life of therapeutic protein: X in an unfused state;
- (4) a nucleic acid molecule (II) comprising a polynucleotide sequence encoding (I);
  - (5) a vector comprising (II); and
  - (6) a host cell comprising (II).

ACTIVITY - Cytostatic; anorectic; immunosuppressive; antidiabetic; antirheumatic; antiarthritic; psoriatic. No supporting data is given.

MECHANISM OF ACTION - None given.

USE - Albumin fusion proteins are stabilized therapeutic proteins e.g. antibodies to C5, C242 and CD80 useful for treating various diseases and disorders such as non-Hodgkin's lymphoma, cancer, obesity, transplant rejection, type I diabetes mellitus, rheumatoid arthritis and psoriasis.

ADVANTAGE - Fusing albumin to therapeutic proteins stabilizes the therapeutic protein, extends the shelf life and retains the in vitro or in vivo biological activity. It also reduces the need to formulate protein solutions with large excesses of carrier proteins to prevent loss of therapeutic proteins due to factors such as binding to the container. The fusion proteins are easily dispensed with a simple formulation requiring minimal post storage manipulation.

The fusion of therapeutic proteins to albumin confers stability in aqueous or other solution. A solution of 200 microgram/ml of human albumin (HA)-human growth hormone (hGH) was prepared in tissue culture media containing 5% horse serum and the solution incubated at 37 degrees C starting at time zero. A sample was removed and tested for its biological activity in the Nb2 cell assay at 2 ng/ml final concentration. The biological activity of HA-gHG remained essentially intact after 5 weeks of incubation at 37 degrees C. The recombinant hGH used as control lost its biological activity in the first week of the experiment.

Dwg.0/20

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FΑ AB; DCN

CPI: B04-B04D4; B04-E02F; B04-E03A; B04-E08; B04-F0100E; B04-G01; B04-N02B0E; B04-P0100E; B11-C07A; B12-K04A; B14-C09B; B14-E12; B14-G02C; B14-H01; B14-N17C; B14-S04; D05-H11; D05-H12A; D05-H12C; D05-H12E; D05-H14; D05-H16; D05-H17C; D05-H17C1 TECH UPTX: 20011129

TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Protein: The albumin activity is the ability to prolong the shelf life of the therapeutic protein: X compared to the shelf life of therapeutic protein: X in the unfused state. (I) has a greater shelf life than the therapeutic protein: X in the unfused state. The in vitro or in vivo biological activity of (I) is greater than the in vitro or in vivo activity of therapeutic protein:  $\boldsymbol{X}$ or its fragment or variant in an unfused state. (I) comprises 2 therapeutic protein: X or their fragments or variants, which are different from each other. Therapeutic protein: X or its fragment or variant is separated from the albumin or its fragment or variant by a linker. (I) comprises a therapeutic protein: X or its fragment or variant I-inserted into an albumin comprising amino acids 54-61, 76-89, 92-100, 170-176, 247-252, 266-277, 280-288, 362-368, 439-447, 462-475, 478-486 or 560-566 of S18. (I) further comprises a secretion leader sequence. (I) has the formula: R1-L-R2; R2-L-R1; or R1-L-R2-L-R1, where:

R1 = therapeutic protein: X or its fragment or variant;
L = peptide linker; and

R2 = albumin comprising S18.

ABEX

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(I) is non-glycosylated and expressed in a glycosylation and protease deficient yeast cell. Alternatively (I) is expressed in a mammalian cell in culture.

Preferred Method: The disease or disorder comprises indication: Y. Preparation: (I) are prepared by standard recombinant techniques. UPTX: 20011129

WIDER DISCLOSURE - Also disclosed as new are:

- (1) transgenic organisms modified to contain (II) to express (I);
- (2) antibodies that bind to a therapeutic protein;
- (3) generating antibodies that bind to a therapeutic protein;
- (4) polynucleotides encoding the antibody;
- (5) diagnosing a disorder comprising assaying the expression of the therapeutic protein in cells or body fluid of an individual using antibodies specific to the therapeutic protein and comparing the level of gene expression with a standard gene expression level, where an increase or decrease in the assayed gene expression level is indicative of a particular disorder; and
- (6) a diagnostic kit for use in screening serum containing antigens of a therapeutic protein comprising an antibody immunoreactive with the antigen.

ADMINISTRATION - 0.1-100 mg/kg of body weight, preferably 1-10 mg/kg of body weight of antibodies are administered by standard routes.

EXAMPLE - Preparation of human albumin fusion proteins was as follows. The cDNA for interferon (IFN) alpha was isolated from cDNA libraries by reverse transcription-polymerase chain reaction (PCR) and by PCR using a series of overlapping synthetic oligonucleotides primers using standard methods. The cDNA was tailored at the 5' and 3' ends to generate restriction sites so that oligonucleotide linkers could be used to clone the cDNA into a vector containing the cDNA for human albumin (HA). This could be at the N or C terminus of the HA sequence with(out) use of a spacer sequence. The IFN alpha cDNA was cloned into a vector such as pPPC0005 from which the complete expression cassette was excised and inserted into the plasmid pSAC35 to allow the expression of the albumin fusion protein in yeast. The albumin fusion protein was collected and purified from the media and tested for its biological activity.

- L88 ANSWER 5 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN
  - AN **2001-602931** [68] WPIX
  - CR 2001-611723 [70]; 2001-616754 [71]; 2001-616755 [71]; 2001-616756 [71]; 2002-010886 [01]; 2002-179329 [23]; 2003-810996 [76]; 2004-033644 [03]

DNC C2001-178694

- TI Albumin fusion proteins comprising a therapeutic protein and albumin, useful in the treating metastatic renal cell carcinoma, metastatic melanoma, malignant melanoma, renal cell carcinoma, HIV (human immunodeficiency virus) or infection.
- DC B04 D16
- IN PRIOR, C P; ROSEN, C A; SADEGHI, H; TURNER, A J
- PA (HUMA-N) HUMAN GENOME SCI INC; (PRIN-N) PRINCIPIA PHARM CORP; (PRIO-I) PRIOR C P; (ROSE-I) ROSEN C A; (SADE-I) SADEGHI H; (TURN-I) TURNER A J CYC 96
- PI WO 2001079258 A1 20011025 (200168)\* EN 325p C07K001-00
  - RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZW
  - W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

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AU 2001059066 A 20011030 (200219)
                                                     C07K001-00
                  A1 20030115 (200313) EN
                                                    C07K001-00
         R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
            RO SE SI TR
     US 2003171267 A1 20030911 (200367)
                                                     A61K038-38
     JP 2003530838 W 20031021 (200373) 430p
                                                    C12N015-09
    WO 2001079258 A1 WO 2001-US12008 20010412; AU 2001059066 A AU 2001-59066
ADT
     20010412; EP 1274720 A1 EP 2001-932549 20010412, WO 2001-US12008 20010412;
     US 2003171267 A1 Provisional US 2000-229358P 20000412, Provisional US
     2000-199384P 20000425, Provisional US 2000-256931P 20001221, US
     2001-833117 20010412; JP 2003530838 W JP 2001-576855 20010412, WO
     2001-US12008 20010412
    AU 2001059066 A Based on WO 2001079258; EP 1274720 A1 Based on WO
     2001079258; JP 2003530838 W Based on WO 2001079258
PRAI US 2000-256931P 20001221; US 2000-229358P 20000412; US 2000-199384P
     20000425; US 2001-833117
                                20010412
     ICM A61K038-38; C07K001-00; C12N015-09
         A01N037-18; A61K035-12; A61K035-76; A61K038-00; A61K038-21;
          A61K038-22; A61K038-23; A61K038-27; A61K047-48; A61K048-00;
          A61P001-04; A61P003-10; A61P003-14; A61P005-10; A61P009-10;
          A61P015-08; A61P017-00; A61P017-02; A61P017-06; A61P017-14;
          A61P019-00; A61P019-02; A61P019-08; A61P019-10; A61P021-00;
         A61P025-00; A61P025-02; A61P025-28; A61P029-00; A61P031-14;
         A61P031-18; A61P031-20; A61P035-00; A61P035-02; A61P035-04;
          A61P037-00; A61P037-06; C07K014-55; C07K014-565; C07K014-585;
          C07K014-60; C07K014-62; C07K014-635; C07K014-76; C07K014-765;
          C07K019-00; C12N001-19; C12N005-10
AΒ
    WO 200179258 A UPAB: 20040112
```

NOVELTY - Albumin fusion proteins (P1) comprising a therapeutic protein (T1) (or its fragment or variant having the activity of T1) and albumin comprising the 585 amino acid sequence (I) defined in the specification (or its fragment or variant having albumin activity), are new.

 ${\tt DETAILED}$  <code>DESCRIPTION</code> - <code>INDEPENDENT</code> <code>CLAIMS</code> are also included for the following:

- (1) a kit comprising a composition containing P1;
- (2) a method of treating a disease or disorder, preferably modulated by T1, in a patient, comprising administering P1;
- (3) a method of extending the shelf-life of T1, comprising fusing T1 or its fragment or variant, to **albumin** or its fragment or variant, where the shelf-life of T1 or its fragment or variant as part of a fused protein is extended when compared to T1 or its fragment or variant in an unfused state;
  - (4) a nucleic acid (N1) comprising a nucleotide sequence encoding P1;
  - (5) a vector comprising N1; and
  - (6) a host cell comprising N1.

ور والمجترية

ACTIVITY - Cytostatic; antiviral; antiinflammatory; antileukemic; antiarthritic; antirheumatic; immunosuppressive; antidiabetic; cardiant; nootropic; neuroprotective; antimicrobial; vulnerary.

To test whether sympathetic neuronal cell viability is supported by an albumin fusion protein, the chicken embryo neuronal survival assay (Senaldi, et al., Proc. Natl. Acad., Sci., U.S.A, 96:11458-63 (1998)). Briefly, motor and sympathetic neurons were isolated from chicken embryos, resuspended in L15 medium (with 10% fetal calf serum (FCS), glucose, sodium selenite, progesterone, conalbumin, putrescine and insulin) and Dulbecco's modified Eagles medium (with 10% FCS, glutamine, penicillin, and 25 mM Hepes buffer (pH 7.2)), respectively and incubated at 37 degrees Centigrade in 5% carbon-dioxide in the presence of different concentrations of the purified fusion protein, as well as negative control lacking any cytokine, After 3 days, neuronal survival was determined by evaluation of cellular morphology, and through the use of the colorimetric assay of Mosmann (Mosmann, T., J. Immunol., Methods, 65:55-63 (1983)). Enhanced neuronal cell viability as compared to the

controls lacking cytokine is indicative of the ability of the albumin fusion protein to enhance the survival of neuronal cells. MECHANISM OF ACTION - Gene therapy.

USE - When the therapeutic protein, or its fragment or variant is IL-2, P1 is used to treat metastatic renal cell carcinoma, metastatic melanoma, malignant melanoma, renal cell carcinoma, HIV (human immunodeficiency virus) infection, inflammatory bowel disorder, Kaposi's sarcoma, leukemia, multiple sclerosis, rheumatoid arthritis, transplant rejection, type 1 diabetes mellitus, lung cancer, acute myeloid leukemia, hepatitis C, non-hodgkin's lymphoma or ovarian cancer (claimed).

The albumin fusion proteins are also useful in the treatment, prevention, diagnosis, and/or detection of diseases, disorders such as immune system disorders (e.g. transplant rejection), blood related disorders (e.g. myocardial infarction), hyperproliferative disorders (e.g. childhood acute myeloid leukemia), renal disorders (e.g. glomerulonephritis), cardiovascular disorders (e.g. arrhythmias), respiratory disorders (e.g. non-allergic rhinitis), neurological diseases (e.g. Alzheimer's disease), endocrine disorders (e.g. pheocytochroma), reproductive system disorders (e.g. syphilis), infectious diseases (e.g. measles), gastrointestinal disorders (e.g. irritable bowel syndrome) and wound healing.

Dwg.0/14

FS CPI

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AB; DCN FΑ

MC CPI: B04-C01; B04-E02F; B04-E08; B04-F0100E; B04-F1100E;

B04-H05; B04-H06; B04-J04; B04-N0200E;

B04-N02A0E; B14-A02B1; B14-C09B; B14-D01; B14-E10C; B14-F01; B14-F02; B14-G02; B14-H01; B14-J01; B14-K01; B14-N10; B14-N12;

B14-N14; B14-N17B; B14-S01; B14-S03; B14-S04; **D05-H12B2**;

D05-H12E; D05-H14

TECH

UPTX: 20011121 TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Fusion Protein: The albumin activity is the ability to prolong the shelf-life of T1 compared to the shelf-life of T1 in an unfused state. The albumin fragment or variant comprises amino acids 1-387 of (I). T1 comprises interleukin 2 (IL-2). The T1 fragment or variant has T cell proliferative activity or T cell activation activity. Tl or its fragment or variant, comprises a protein selected from calcitonin, growth hormone releasing factor, IL-2 fusion protein, insulin-like growth factor-1, interferon beta or parathyroid hormone. T1 or its fragment or variant is fused to the C-terminal of the albumin or the C-terminus of the fragment or variant of albumin. Alternatively, T1 or its fragment or variant is fused to the N-terminal of the albumin or the N-terminus of the fragment or variant of albumin. Alternatively, T1 or its fragment or variant is fused to the N-terminus and C-terminus of the albumin, or the N-terminus and C-terminus of the fragment or variant of albumin. P1 comprises a first T1 or its fragment or variant, and a second T1 or its fragment or variant, where the first T1 is different from the second T1. T1 or its fragment or variant is separated from the albumin or the fragment or variant of albumin by a linker. Preferably, P1 is of the formula (S1), (S2) or (S3). R1-L-R2 (S1); R2-L-R1 (S2); or R1-L-R2-L-R1 (S3). R1 = is T1 or its fragment or variant; L = is a peptide linker; and

R2 = is albumin comprising the sequence of (I), or its fragment

The shelf-life of the albumin fusion protein is greater than the shelf-life of T1 or its fragment or variant in an unfused state. The in vitro or in vivo biological activity of Tl or its fragment or variant, fused to albumin or its fragment or variant, is greater than the in vitro or in vivo, respectively, biological activity of T1 or its fragment or variant, in an unfused state.

Alternatively, P1 comprises T1 or its fragment or variant, inserted into an albumin comprising the sequence of (I) or its fragment or variant. Preferably, the albumin comprises residues 54-61, 76-89, 92-100, 170-176, 247-252, 266-277, 280-288, 362-368, 439-447, 462-475, 478-486, or 560-566 of (I). The portion of albumin is sufficient to prolong the shelf-life and in vitro and in vivo biological activity of T1 or its fragment or variant, as compared to the shelf-life and in vitro and in vivo biological activity of T1 or its fragment or variant, in an unfused state.

P1 is non-glycosylated and expressed in yeast which is glycosylation deficient. The yeast may also be protease deficient. Alternatively, P1 is

expressed by a mammalian cell in culture. Pl further comprises a secretion

leader sequence.
ABEX UPTX: 20011121

ADMINISTRATION - The **albumin** fusion proteins can be administered orally, rectally, parenterally, intracisternally, intravaginally, intraperitoneally, topically, bucally, or as an oral or nasal spray. The dosage is 1 microgram/kg/day to 10 mg/kg/day, preferably 0.01 to 1, mg/kd/day. If given continuously, the **albumin** fusion protein is typically administered at a dose rate of 1-50 micrograms/kg/hour, either by 1-4 injections per day or by continuous subcutaneous infusions.

EXAMPLE - The cDNA for the growth factor of interest such as interferon growth factor 1 (IGF-1) can be isolated using a variety of means including but not exclusively, from cDNA libraries, by reverse transcriptasepolymerase chain reaction (PCR) and by PCR using a series of overlapping synthetic oligonucleotide primers, all using standard methods (see GenBank Acc. Number NP-000609). The cDNA can be tailored at the 5' and 3' ends to generate restriction sites, such that the oligonucleotide linkers can be used, for cloning of the cDNA into a vector containing the cDNA for human serum albumin (HA). This can be a the N or C-terminus with or without the use of a spacer sequence. The growth factor cDNA was cloned into a vector such as pPPC0005, pScCHSA, pScNHSA or pC4:HSA from which the complete expression cassette is then excised and inserted into the plasmid pSAC35 to allow the expression of the albumin fusion protein in yeast. The albumin fusion protein secreted from the yeast can then be collected and purified from the media and tested for its biological activity. For expression in mammalian cell lines a similar procedure is adopted except that the expression cassette used employs a mammalian promoter, leader sequence and terminator. This expression cassette is then excised and inserted into a plasmid suitable for the transfection of mammalian cell lines.

L88 ANSWER 6 OF 6 WPIX COPYRIGHT 2004 THOMSON DERWENT on STN

AN 1996-300388 [30] WPIX

DNC C1996-095415

TI New chimeric proteins for treatment of septic shock, psoriasis, cancers etc. - comprise cytokine bonded to polypeptide which is enzymatically inactive in humans, increases half-life and prevents cytokine(s) from crossing blood brain barrier.

DC B04

4 F. S.

د. وأيتذبه

IN STEELE, A; STROM, T B; ZHENG, X; ZHENG, X X

PA (BETH-N) BETH ISRAEL HOSPITAL ASSOC

CYC 20

PI WO 9618412 A1 19960620 (199630)\* EN 58p A61K038-19 RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE W: CA JP

EP 793504 A1 19970910 (199741) EN A61K038-19

R: CH DE FR GB IT LI SE

JP 11501506 W 19990209 (199916) 49p C12N015-09

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B1 20020611 (200244)
                                                     A61K038-20
     US 6403077
                                                     C07K014-54
     US 6410008
                  B1 20020625 (200246)
     US 2002173628 A1 20021121 (200279)
                                                     A61K038-52
     US 2003026778 A1 20030206 (200318)
                                                     A61K038-20
     WO 9618412 A1 WO 1995-US16046 19951212; EP 793504 A1 EP·1995-943058
     19951212, WO 1995-US16046 19951212; JP 11501506 W WO 1995-US16046
     19951212, JP 1996-519191 19951212; US 6403077 B1 CIP of US 1994-355502
     19941212, Cont of US 1995-431535 19950428, US 1997-968905 19971106; US
     6410008 B1 US 1994-355502 19941212; US 2002173628 A1 Cont of US
     1994-355502 19941212, US 2002-145481 20020514; US 2003026778 A1 CIP of US
     1994-355502 19941212, Cont of US 1997-968905 19971106, US 2002-145517
     20020514
     EP 793504 Al Based on WO 9618412; JP 11501506 W Based on WO 9618412; US
     2002173628 A1 Cont of US 6410008; US 2003026778 A1 Cont of US 6403077, CIP
     of US 6410008
PRAI US 1995-431535
                      19950428; US 1994-355502
                                                 19941212; US 1997-968905
     19971106; US 2002-145481
                                20020514; US 2002-145517
     2.Jnl.Ref; US 5231012
     ICM A61K038-19; A61K038-20; A61K038-52; C07K014-54; C12N015-09
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         A61K038-00; A61K038-21; A61K038-38; A61K039-395;
          C07K014-52; C07K014-525; C07K014-53; C07K014-535;
          C07K014-545; C07K014-55; C07K014-555; C07K014-76;
          C07K014-765; C07K016-18; C07K016-46; C07K019-00;
          C12N009-10; C12N015-02; C12N015-24; C12P021-02
AB
          9618412 A UPAB: 19960731
     Chimeric protein comprises a cytokine bonded to a polypeptide which is
     enzymatically inactive in humans and which increases the circulating
     half-life of the cytokine in vivo by a factor of 1.
           Also claimed is the use of interleukin-10 (IL-10)/Fc in the preparation
     of a medicament for inhibiting granuloma formation in a patient.
          USE - The chimeric proteins can be used to treat conditions for which
     the corresp. cytokines are used, e.g. septic shock, granulomatous
     disorders (e.g. schistosomiasis), multiple sclerosis, psoriasis,
     rheumatoid arthritis, cancers and virus infections. Chimeric proteins
     including a lytic Fc region can also be used to deplete patients of
     suppressor lymphocytes and to treat chronic infections such as those
     associated with suppression of the immune system.
          ADVANTAGE - The enzymatically inactive polypeptides extend the
     circulating half-life of the cytokines in vivo by a factor of 10
     (claimed). In addition, they can prevent the cytokines from crossing the
     blood brain barrier and causing adverse side effects.
     Dwg.0/15
     CPI
FS
FΑ
     AΒ
MC
     CPI: B04-B04; B04-G01; B04-H02; B04-H04A; B04-H04C; B04-H08;
          B04-N02; B14-A01; B14-C09B; B14-N17C; B14-S01; B14-S06
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                E ALBUMIN/CT
L1
            753 S E3
            132 S E11
L2
                E E47+ALL
          80101 S E2+NT
L3
                E E33+ALL
            566 S E3, E2
L4
          25218 S E2+NT
L5
         157881 S ?ALBUMIN?
L6
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و المجالة

٠٠ ويُعتبر

وروانيتون

مه وفيتقاله

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181833 S L1-L6
L7
L8
            2969 S BDNF OR BD NF
            2881 S BRAIN DERIVED NEUROTROPHIC FACTOR
L9
            2883 S (BD OR BRAIN DERIVED) () (NF OR NEUROTROPHIC FACTOR)
L10
                 E NEUROTROPHIC FACTOR/CT
L11
            141 S E10
            2554 S E26
                 E E25+ALL
             789 S E3-E5 AND BRAIN DERIVED
L13
L14
            679 S E12,E13
L15
            3242 S E2+NT (L) BRAIN DERIVED
L16
              64 S L7 AND L8-L15
           19234 S INTERFERONALPHA OR ALPHAINTERFERON OR INTERFERONBETA OR BETAI
L17
                 E INTERFERON/CT
· L18
             302 S E3-E19
L19
           18390 S E85-E101
                 E INTERFERONS/CT
                 E E3+ALL
           18391 S E7, E6 (L) (ALPHA OR BETA)
             546 S L7 AND L17-L20
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      FILE 'HCAPLUS' ENTERED AT 15:30:37 ON 02 FEB 2004
 L24
            2026 S L23
L25
             859 S TISSUE INHIBITOR (1W) METALLOPROTEINASE 1
L26
              27 S METALLOPROTEINASE INHIBITOR 1
L27
             651 S TIMP1
              12 S FIBROBLAST COLLAGENASE INHIBITOR
L28
L29
             91 S L7 AND L22, L24-L28
L30
            678 S L16, L21, L29
            9815 S IFNALPHA OR IFNBETA OR ALPHAIFN OR BETAIFN OR IFN(A) (ALPHA OR
L31
L32 ·
             119 S L7 AND L31
L33
             700 S L30, L32
              62 S L33 AND (FUSION OR FUSE OR FUSED OR FUSES OR FUSING)
L34
L35
             167 S L33 AND RECOMBIN?
L36
              44 S L33 AND CHIMER?
L37
             202 S L34-L36
                 E ROSEN C/AU
 L38
              27 S E3, E4
                 E ROSEN CRAIG/AU
L39
             625 S E3-E5
                 E HASELTINE W/AU
             302 S E3, E4, E7-E10
L40
              10 S L33 AND L38-L40
L41
                 E HUMAN GENOME SCI/PA, CS
L42
             975 S E5-E37
L43
              13 S L33 AND L42
              13 S L41, L43
L44
              13 S L44 AND L37
L45
L46
               9 S L45 AND (SHELFLIFE OR SHELF LIFE)
L47
               4 S L45 NOT L46
                 SEL DN AN 1 4
L48
               2 S L47 NOT E1-E6
              11 S L46, L48
L49
                 SEL RN
                 DEL SEL
                 E FUSION PROTEIN/CT
 L50 ·
           11933 S E9
                 E E9+ALL
            3795 S E3, E4
 L51
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5 S L51 AND L33
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L53
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L54
L55
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L56
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L57
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L59
             7 S L58 AND L29 '
L60
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L62
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L64
L65
             29 S L64 AND ?ALBUMIN?
             29 S L64 AND (INF? OR INTERFERON OR TIMP? OR NEUROTROPHIC?)
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L67
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L68
           1564 S L17/BIX OR LL31/BIX
L69
            80 S L22/BIX OR L25/BIX OR L26/BIX OR L27/BIX OR L28/BIX
L70
L71
            124 S L67 AND L68-L70
          11209 S ?ALBUMEN?/BIX OR L67
L72
            513 S (A61K038-38 OR C07K014-76 OR C07K014-765 OR C12N015-14)/IC,IC
L73
L74
          11377 S L72,L73
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           216 S L74 AND L75
L77
            111 S L74 AND L76
L78
            129 S L74 AND L68, L69, L70
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L80
            311 S L77-L79
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L81
           7242 S (D05-H12B OR D05-H12B2)/MC
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     FILE 'HCAPLUS' ENTERED AT 16:25:16 ON 02 FEB 2004
     FILE 'REGISTRY' ENTERED AT 16:26:59 ON 02 FEB 2004
L89
              1 S 507485-69-0
L90
              1 S 472960-22-8
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=>

-52.5

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NEWS
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                 CA/CAplus records now contain indexing from 1907 to the
NEWS
         SEP 09
                 present
                 INPADOC: Legal Status data reloaded
NEWS
         DEC 08
                 DISSABS now available on STN
NEWS
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         SEP 29
NEWS
     6
         OCT 10
                 PCTFULL: Two new display fields added
                 BIOSIS file reloaded and enhanced
NEWS
      7
         OCT 21
                 BIOSIS file segment of TOXCENTER reloaded and enhanced
         OCT 28
NEWS
     8
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NEWS
     9
         DEC 08
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NEWS 10
         DEC 08
                 IMS file names changed
NEWS 11
                 Experimental property data collected by CAS now available
NEWS 12
         DEC 09
                 in REGISTRY
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         DEC 09
NEWS 13
                 DGENE: Two new display fields added
         DEC 17
NEWS 14
NEWS 15
         DEC 18
                 BIOTECHNO no longer updated
                 CROPU no longer updated; subscriber discount no longer
         DEC 19
NEWS 16
                 available
                 Additional INPI reactions and pre-1907 documents added to CAS
         DEC 22
NEWS 17
                 databases
                 IFIPAT/IFIUDB/IFICDB reloaded with new data and search fields
         DEC 22
NEWS 18
                 ABI-INFORM now available on STN
NEWS 19
         DEC 22
                 Source of Registration (SR) information in REGISTRY updated
NEWS 20
         JAN 27
                 and searchable
                 A new search aid, the Company Name Thesaurus, available in
         JAN 27
NEWS 21
                 CA/CAplus
                 German (DE) application and patent publication number format
NEWS 22
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                 changes
              DECEMBER 28 CURRENT WINDOWS VERSION IS V7.00, CURRENT
NEWS EXPRESS
              MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP),
              AND CURRENT DISCOVER FILE IS DATED 23 SEPTEMBER 2003
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FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004

=> file medline, uspatful, dgene, embase, wpids

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 13:52:47 ON 06 FEB 2004

FILE 'USPATFULL' ENTERED AT 13:52:47 ON 06 FEB 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'DGENE' ENTERED AT 13:52:47 ON 06 FEB 2004 COPYRIGHT (C) 2004 THOMSON DERWENT

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FILE 'WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004 COPYRIGHT (C) 2004 THOMSON DERWENT

=> s albumin fusion proteins

2835 ALBUMIN FUSION PROTEINS

=> s cerebus protein

1 CEREBUS PROTEIN

=> s 11 and 12

0 L1 AND L2

=> s (cerebus protein) and albumin

0 (CEREBUS PROTEIN) AND ALBUMIN

=> s 12 and fusion

0 L2 AND FUSION

=> d 12 ti abs ibib tot

ANSWER 1 OF 1 WPIDS COPYRIGHT 2004 THOMSON DERWENT on STN **L2** 

TI Human and murine cerebus-like proteins - used for treating tissue defects and degenerative nerve conditions.

WPIDS ΔN 1999-106054 [09]

2003-298696 [29] CB

AB 9901553 A UPAB: 20030505

A novel isolated DNA sequence comprises a DNA sequence selected from: (a) nucleotides beginning at # 1, 52, 55, 58, 61, 64, 67, 70, 73, 121, 256, 259, 262, 265, 268, 171, or 484 and ending at # 723 or 801 of the 804 bp DNA sequence given in the specification; and (b) sequences which hybridise to (a) under stringent hybridisation conditions and encode a protein which exhibits cerebus activity. Also claimed are: (1) an isolated DNA sequence comprising nucleotides encoding amino acids beginning at #1, 18 to 25, 41, 85 to 91 or 152, and ending at #241 or 267 of the 267 amino acid sequence given in the specification; (2) a vector comprising either of the above DNA molecules in operative association with an expression control sequence; (3) an isolated DNA molecule comprising nucleotides 268-801 of the 272 amino acid sequence given in the specification (sic), or naturally occurring allelic sequences of it; (4) a vector comprising the DNA of (4) in operative association with an expression control sequence; (5) an isolated DNA molecule encoding mammalian cerebus protein , comprising nucleotides 268-801 of the 804 bp DNA sequence given in the specification; (6) a vector comprising the DNA of (5) in operative

association with an expression control sequence; (7) a host cell

transformed with the vector of (2), (4) or (6); (8) a purified mammalian cerebus protein comprising the 267 amino acid sequence given in the specification; (9) a purified mammalian cerebus protein comprising residues 90-267 of the 272 amino acid sequence given in the specification; and (10) antibodies to the cerebus protein of (8) or (9).

USE - The host cell of (7) can be used to produce the mammalian cerebus proteins (claimed). Compositions containing the protein can be used in the formation of neurons and related neural cells and tissues, such as Schwann cells, glial cells, and astrocytes, as well as liver, pancreas, lung, heart, kidney, spleen, stomach, and cardiac tissue and cells. They may also be used to treat precursor or stem cells. The compositions can also be used for treating tissue defects, and healing and maintenance of various types of tissues and wounds. The mammalian cerebus protein containing compositions may also be used to treat or prevent degenerate nerve conditions such as Parkinson's disease, Alzheimer's disease, and Lou Gehrig's disease. They can also be used to treat osteoporosis, rheumatoid arthritis, osteoarthritis, and other abnormalities of connective tissue, or of other organs or tissues.

ACCESSION NUMBER:

1999-106054 [09] WPIDS

CROSS REFERENCE:

2003-298696 [29]

DOC. NO. CPI:

C1999-031758

TITLE:

Human and murine cerebus-like proteins - used for treating tissue defects and degenerative nerve

conditions.

DERWENT CLASS:

B04 D16

INVENTOR(S):

DEROBERTIS, E M; FOLLETTIE, M

PATENT ASSIGNEE(S):

(GEMY) GENETICS INST INC; (REGC) UNIV CALIFORNIA

COUNTRY COUNT:

PATENT INFORMATION:

| PATENT NO | KIND DATE | WEEK | LA PG |
|-----------|-----------|------|-------|
|           |           |      |       |

A1 19990114 (199909)\* EN 50 WO 9901553

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG

57

UZ VN YU ZW

A 19990125 (199923) AU 9878140

US 5935852 A 19990810 (199938)

EP 1012278 A1 20000628 (200035) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

MX 2000000242 A1 20010601 (200235)

JP 2002511762 W 20020416 (200242)

B 20020620 (200252) AU 749031

#### APPLICATION DETAILS:

| PAT | ENT NO K   | IND | API | PLICATION    | DATE     |
|-----|------------|-----|-----|--------------|----------|
| WO  | 9901553    | A1  | wo  | 1998-US11462 | 19980603 |
| ΑU  | 9878140    | A   | ΑU  | 1998-78140   | 19980603 |
| US  | 5935852    | A   | US  | 1997-887997  | 19970703 |
| ΕP  | 1012278    | A1  | EΡ  | 1998-926263  | 19980603 |
|     |            |     | WO  | 1998-US11462 | 19980603 |
| MX  | 2000000242 | A1  | MΧ  | 2000-242     | 20000105 |
| JP  | 2002511762 | W   | WO  | 1998-US11462 | 19980603 |
|     |            |     | JP  | 1999-507147  | 19980603 |
| ΑU  | 749031     | В   | ΑU  | 1998-78140   | 19980603 |
|     |            |     |     |              |          |

### FILING DETAILS:

| PATENT N                                      | O KIND           |  | :     | PATENT   | NO                      |
|---|------------------|--|-------|--|-------------------------|
| AU 98781/<br>EP 10122<br>JP 20025<br>AU 74903 | 78 A1<br>11762 W | Based on<br>Based on<br>Based on<br>Previous<br>Based on | Publ. | WO 990<br>WO 990<br>WO 990<br>AU 987<br>WO 990 | 01553<br>01553<br>78140 |

PRIORITY APPLN. INFO: US 1997-887997 19970703

=> file biosis

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 20.32 20.53

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004 COPYRIGHT (C) 2004 BIOLOGICAL ABSTRACTS INC.(R)

FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 4 February 2004 (20040204/ED)

FILE RELOADED: 19 October 2003.

=> d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN

L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

=> s 12

0 CEREBUS

1361492 PROTEIN

L6 0 CEREBUS PROTEIN

(CEREBUS (W) PROTEIN)

=> file medline, uspatful, dgene, embase, wpids, biosis, japio, fsta, jicst COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST 0.85 21.38

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FILE 'USPATFULL' ENTERED AT 14:00:26 ON 06 FEB 2004
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FILE 'JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004 COPYRIGHT (C) 2004 Japan Science and Technology Agency (JST)

=> s 12

L7 1 L2

=> d 17 ti abs ibib tot

L7 ANSWER 1 OF 1 WPIDS COPYRIGHT 2004 THOMSON DERWENT on STN TI Human and murine cerebus-like proteins - used for treating tissue defects

and degenerative nerve conditions.

AN 1999-106054 [09] WPIDS

CR 2003-298696 [29]

AB WO 9901553 A UPAB: 20030505

A novel isolated DNA sequence comprises a DNA sequence selected from: (a) nucleotides beginning at # 1, 52, 55, 58, 61, 64, 67, 70, 73, 121, 256, 259, 262, 265, 268, 171, or 484 and ending at # 723 or 801 of the 804 bp DNA sequence given in the specification; and (b) sequences which hybridise to (a) under stringent hybridisation conditions and encode a protein which exhibits cerebus activity. Also claimed are: (1) an isolated DNA sequence comprising nucleotides encoding amino acids beginning at #1, 18 to 25, 41, 85 to 91 or 152, and ending at #241 or 267 of the 267 amino acid sequence given in the specification; (2) a vector comprising either of the above DNA molecules in operative association with an expression control sequence; (3) an isolated DNA molecule comprising nucleotides 268-801 of the 272 amino acid sequence given in the specification (sic), or naturally occurring allelic sequences of it; (4) a vector comprising the DNA of (4) in operative association with an expression control sequence; (5) an isolated DNA molecule encoding mammalian cerebus protein , comprising nucleotides 268-801 of the 804 bp DNA sequence given in the specification; (6) a vector comprising the DNA of (5) in operative association with an expression control sequence; (7) a host cell transformed with the vector of (2), (4) or (6); (8) a purified mammalian cerebus protein comprising the 267 amino acid sequence given in the specification; (9) a purified mammalian cerebus protein comprising residues 90-267 of the 272 amino acid sequence given in the specification; and (10) antibodies to the cerebus protein of (8) or (9).

USE - The host cell of (7) can be used to produce the mammalian cerebus proteins (claimed). Compositions containing the protein can be used in the formation of neurons and related neural cells and tissues, such as Schwann cells, glial cells, and astrocytes, as well as liver, pancreas, lung, heart, kidney, spleen, stomach, and cardiac tissue and cells. They may also be used to treat precursor or stem cells. The compositions can also be used for treating tissue defects, and healing and maintenance of various types of tissues and wounds. The mammalian cerebus protein containing compositions may also be used to treat or prevent degenerate nerve conditions such as Parkinson's disease, Alzheimer's disease, and Lou Gehrig's disease. They can also be used to treat osteoporosis, rheumatoid arthritis, osteoarthritis, and other abnormalities of connective tissue, or of other organs or tissues. Dwg.0/0

ACCESSION NUMBER: 1999-106054 [09] WPIDS

2003-298696 [29] CROSS REFERENCE: C1999-031758 DOC. NO. CPI:

Human and murine cerebus-like proteins - used for TITLE:

treating tissue defects and degenerative nerve

conditions.

DERWENT CLASS: B04 D16

DEROBERTIS, E M; FOLLETTIE, M INVENTOR(S):

(GEMY) GENETICS INST INC; (REGC) UNIV CALIFORNIA PATENT ASSIGNEE(S):

COUNTRY COUNT:

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LAPG

WO 9901553 A1 19990114 (199909) \* EN 50

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL

OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE GH GM GW HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG

MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG

UZ VN YU ZW

AU 9878140 A 19990125 (199923) US 5935852 A 19990810 (199938)

EP 1012278 A1 20000628 (200035) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

MX 2000000242 A1 20010601 (200235)

JP 2002511762 W 20020416 (200242) 57

AU 749031 B 20020620 (200252)

#### APPLICATION DETAILS:

| PATENT NO K              | IND    | APPLICATION                       | DATE                 |
|--------------------------|--------|-----------------------------------|----------------------|
| WO 9901553<br>AU 9878140 | A1     | WO 1998-US11462<br>AU 1998-78140  | 19980603<br>19980603 |
| US 5935852               | A<br>A | US 1997-887997                    | 19970703             |
| EP 1012278               | A1     | EP 1998-926263<br>WO 1998-US11462 | 19980603<br>19980603 |
| MX 2000000242            | A1     | MX 2000-242                       | 20000105             |
| JP 2002511762            | W      | WO 1998-US11462                   | 19980603             |
|                          |        | JP 1999-507147                    | 19980603             |
| AU 749031                | В      | AU 1998-78140                     | 19980603             |

#### FILING DETAILS:

| PAT | TENT NO K          | IND |                      |       | PAT | TENT NO            |
|-----|--------------------|-----|----------------------|-------|-----|--------------------|
|     | 9878140<br>1012278 |     | Based on<br>Based on |       |     | 9901553<br>9901553 |
| JP  | 2002511762         | W   | Based on             |       | WO  | 9901553.           |
| AU  | 749031             | В   | Previous<br>Based on | Publ. |     | 9878140<br>9901553 |

PRIORITY APPLN. INFO: US 1997-887997 19970703

### => d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

L6 0 S L2

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

L7 1 S L2

=> s TIMP-1 or tissue inhibitor metalloproteinase-1

5 FILES SEARCHED...

8080 TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1

=> s 18 and 11

L9 5 L8 AND L1

=> d 19 ti abs ibib tot

L9 ANSWER 1 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL Albumin fusion proteins

TITLE:
INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2004010134  | A1   | 20040115 |     |
| APPLICATION INFO.:  | US 2001-833245 | A1   | 20010412 | (9) |

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 25 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 2 OF 5 USPATFULL on STN

TI Albumin fusion proteins

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic

acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR (S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003219875  | A1   | 20031127 |     |
| APPLICATION INFO.:  | US 2001-833118 | A1   | 20010412 | (9) |

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

29

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 3 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL

TITLE:
INVENTOR(S):

Albumin fusion proteins

Ballance, David J., Berwyn, PA, UNITED STATES
Sleep, Darrell, West Bridgford, UNITED KINGDOM
Prior, Christopher P., Rosemont, PA, UNITED STATES
Sadeghi, Homayoun, Doylestown, PA, UNITED STATES
Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND   | DATE     |     |
|---------------------|----------------|--------|----------|-----|
| PATENT INFORMATION: | US 2003199043  | <br>А1 | 20031023 |     |
| APPLICATION INFO.:  | US 2001-832501 | A1     | 20031023 | (9) |

NUMBER DATE

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PRIORITY INFORMATION: US 2000-256931P 20001221 (60)

US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility
APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 60

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

14339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 4 OF 5 USPATFULL on STN

TI Albumin fusion proteins

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:244853 USPATFULL Albumin fusion proteins

INVENTOR(S):

TITLE:

Rosen, Craig A., Laytonsville, MD, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003171267  | A1   | 20030911 |     |
| APPLICATION INFO.:  | US 2001-833117 | A1   | 20010412 | (9) |

|          | -            |    | NUMBER                       | DATE                 |   |
|----------|--------------|----|------------------------------|----------------------|---|
| PRIORITY | INFORMATION: |    | 2000-256931P<br>2000-199384P | 20001221<br>20000425 | , |
|          |              | US | 2000-229358P                 | 20000412             | , |

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 59 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 20 Drawing Page(s)

LINE COUNT: 13208

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- L9 ANSWER 5 OF 5 USPATFULL on STN
- TI Albumin fusion proteins
- The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion

proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT. ACCESSION NUMBER: 2003:181414 USPATFULL Albumin fusion proteins TITLE: Rosen, Craig A., Laytonsville, MD, UNITED STATES INVENTOR(S): Haseltine, William A., Washington, DC, UNITED STATES KIND DATE NUMBER \_\_\_\_\_\_\_ A1 20030703 US 200312524.7 PATENT INFORMATION: US 2001-833041 A1 20010412 (9) APPLICATION INFO .: NUMBER DATE \_\_\_\_\_\_ US 2000-256931P 20001221 (60) PRIORITY INFORMATION: 20000425 (60) US 2000-199384P 20000412 (60) US 2000-229358P DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE: ROCKVILLE, MD, 20850 NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: NUMBER OF DRAWINGS: 20 Drawing Page(s) LINE COUNT: 15235 CAS INDEXING IS AVAILABLE FOR THIS PATENT. => d his (FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004) FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004 2835 S ALBUMIN FUSION PROTEINS L11 S CEREBUS PROTEIN L20 S L1 AND L2 L3 0 S (CEREBUS PROTEIN) AND ALBUMIN L40 S L2 AND FUSION L5 FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004 0 S L2 L<sub>6</sub> FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004 1 S L2 L78080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1 L8 5 S L8 AND L1 L9 => s 18 and fusion 378 L8 AND FUSION L10=> s 110 and albumin 221 L10 AND ALBUMIN L11

=> d l12 ti abs ibib tot

L12 ANSWER 1 OF 5 USPATFULL on STN

TI Albumin fusion proteins

=> s 111 and albumin fragment

AB The present invention encompasses albumin fusion

5 L11 AND ALBUMIN FRAGMENT

proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL

TTTLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

| •                   | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2004010134  | A1   | 20040115 |     |
| APPLICATION INFO.:  | US 2001-833245 | A1   | 20010412 | (9) |
|                     |                |      |          |     |

NUMBER DATE \_\_\_\_\_\_ PRIORITY INFORMATION: US 2000-256931P 20001221 (60) US 2000-229358P 20000412 (60)
Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

29

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 2 OF 5 USPATFULL on STN

Albumin fusion proteins ΤI

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                     |    | NUMBER      | KIND      | DATE     |     |
|---------------------|----|-------------|-----------|----------|-----|
|                     |    |             |           |          |     |
| PATENT INFORMATION: | US | 2003219875  | A1        | 20031127 |     |
| APPLICATION INFO.:  | US | 2001-833118 | <b>A1</b> | 20010412 | (9) |

NUMBER DATE PRIORITY INFORMATION: US 2000-256931P 20001221 (60)

US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

# L12 ANSWER 3 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL

TITLE:
INVENTOR(S):

Albumin fusion proteins Ballance, David J., Berwyn, PA, UNITED STATES

Sleep, Darrell, West Bridgford, UNITED KINGDOM Prior, Christopher P., Rosemont, PA, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND      | DATE     |     |
|---------------------|----------------|-----------|----------|-----|
|                     |                |           |          |     |
| PATENT INFORMATION: | US 2003199043  | <b>A1</b> | 20031023 |     |
| APPLICATION INFO.:  | US 2001-832501 | A1        | 20010412 | (9) |

NUMBER DATE
-----PRIORITY INFORMATION: US 2000-256931P 20001221 (60)
US 2000-199384P 20000425 (60)
US 2000-229358P 20000412 (60)

albumin fusion proteins of the invention.

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 60 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 14339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

### L12 ANSWER 4 OF 5 USPATFULL on STN

TI Albumin fusion proteins

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using

these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:2448

2003:244853 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003171267  | A1   | 20030911 |     |
| APPLICATION INFO.:  | US 2001-833117 | A1   | 20010412 | (9) |

NUMBER DATE

PRIORITY INFORMATION:

US 2000-256931P 20001221 (60) US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

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NUMBER OF CLAIMS: EXEMPLARY CLAIM:

59 1

NUMBER OF DRAWINGS:

20 Drawing Page(s)

LINE COUNT:

13208

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 5 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:181414 USPATFULL Albumin fusion proteins

INVENTOR (S):

TITLE:

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Haseltine, William A., Washington, DC, UNITED STATES

|  | NUMBER                          | KIND     | DATE     |     |
|--|---------------------------------|----------|----------|-----|
| PATENT INFORMATION: APPLICATION INFO.: | US 2003125247<br>US 2001-833041 | A1<br>A1 | 20030703 | (9) |

|          |              |    | NUMBER                                       | DATE                             |      |
|----------|--------------|----|--|----------------------------------|------|
| PRIORITY | INFORMATION: | US | 2000-256931P<br>2000-199384P<br>2000-229358P | 20001221<br>20000425<br>20000412 | (60) |

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

29

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

20 Drawing Page(s)

LINE COUNT:

15235

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### => d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN

L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

L6 0 S L2

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

L7 1 S L2

L8 8080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1

L9 5 S L8 AND L1 L10 378 S L8 AND FUSION L11 221 S L10 AND ALBUMIN

L12 5 S L11 AND ALBUMIN FRAGMENT

=> s lll and shelf-life

L13 9 L11 AND SHELF-LIFE

#### => d l13 ti abs ibib tot

L13 ANSWER 1 OF 9 USPATFULL on STN

TI Biospecific contrast agents

Methods and apparatuses for detecting a condition of a sample (including cervical cancers and pre-cancers) through reflectance and/or fluorescence imaging. A sample is obtained. One or more metallic nanoparticles and/or one or more quantum dots are obtained. The one or more metallic nanoparticles and/or one or more quantum dots are coupled to one or more biomarkers of the sample that are associated with the condition. A reflectance and/or fluorescence image of the sample is then taken. The image(s) exhibit characteristic optical scattering from the one or more metallic nanoparticles and/or characteristic fluorescence excitation from the one or more quantum dots to signal the presence of the one or more biomarkers. In this way, the condition can be readily screened or diagnosed.

ACCESSION NUMBER:

2004:31276 USPATFULL

TITLE:

Biospecific contrast agents

INVENTOR (S):

Sokolov, Konstantin, Austin, TX, UNITED STATES Korgel, Brian A., Round Rock, TX, UNITED STATES Ellington, Andrew D., Austin, TX, UNITED STATES Richards-Kortum, Rebecca, Austin, TX, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION: US 2004023415

US 2004023415 A1 20040205

APPLICATION INFO.: US 2003-382136 A1 20030305 (10)

NUMBER DATE

PRIORITY INFORMATION:

US 2002-361924P 20020305 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

Michael C. Barrett, Esq., FULBRIGHT & JAWORSKI, L.L.P.,

600 Congress Avenue, Suite 2400, Austin, TX, 78701

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 44

NUMBER OF DRAWINGS:

11 Drawing Page(s)

LINE COUNT:

3948

### L13 ANSWER 2 OF 9 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL Albumin fusion proteins

TITLE: INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2004010134  | A1   | 20040115 |     |
| APPLICATION INFO.:  | US 2001-833245 | A1   | 20010412 | (9) |

NUMBER DATE

PRIORITY INFORMATION: US 2000-256931P 20001221 (60) US 2000-199384P 20000425 (60)

US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

### L13 ANSWER 3 OF 9 USPATFULL on STN

TI Nanoporous particle with a retained target

AB Porous nanostructured materials, such as porous nanostructured liquid and liquid crystalline particles or materials, incorporate a target substantially within the material which selectively binds a chemical of interest which can diffusion within the porous nanostructured material and be bound by the target. The porous nanostructured materials can be dispersed as particles in a medium in which said chemical of interest is located with low turbidity. Markers which detect binding of said

chemical of interest can be maintained in the medium separate and apart from the target, and any active compound (e.g., an enzyme) associated therewith by the porous nanostructured material, such that detectable changes in the marker only result when the active compounds diffuse out of the porous nanostructured materials after the chemical of interest binds to the target.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:330129 USPATFULL

TITLE:

Nanoporous particle with a retained target

INVENTOR(S):

Anderson, David, Colonial Heights, VA, UNITED STATES

NUMBER KIND DATE -----US 2003232340 A1 20031218 PATENT INFORMATION:

APPLICATION INFO.:

US 2002-170214

A1 20020613

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

WHITHAM, CURTIS & CHRISTOFFERSON, P.C., 11491 SUNSET

HILLS ROAD, SUITE 340, RESTON, VA, 20190

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

119 1

NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

2555

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 4 OF 9 USPATFULL on STN

TI Albumin fusion proteins

AB

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL Albumin fusion proteins

INVENTOR(S):

TITLE:

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

NUMBER KIND DATE -----US 2003219875 A1 US 2001-833118 A1 20031127 PATENT INFORMATION: APPLICATION INFO.: 20010412 (9)

DATE NUMBER

US 2000-256931P 20001221 (60) PRIORITY INFORMATION: US 2000-199384P 20000425 (60)

albumin fusion proteins of the invention.

US 2000-229358P 20000412 (60)

\_\_\_\_\_\_

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

29

18 Drawing Page(s) NUMBER OF DRAWINGS:

LINE COUNT:

15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 5 OF 9 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Ballance, David J., Berwyn, PA, UNITED STATES Sleep, Darrell, West Bridgford, UNITED KINGDOM Prior, Christopher P., Rosemont, PA, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

| ·                                      | NUMBER                          | KIND | DATE                 |     |
|--|---------------------------------|------|----------------------|-----|
| PATENT INFORMATION: APPLICATION INFO.: | US 2003199043<br>US 2001-832501 | A1   | 20031023<br>20010412 | (9) |

|          | •            |    | NUMBER       | DATE     |      |
|----------|--------------|----|--------------|----------|------|
|          |              |    | <del></del>  | ,        |      |
| PRIORITY | INFORMATION: | US | 2000-256931P | 20001221 | (60) |
|          |              | US | 2000-199384P | 20000425 | (60) |
|          |              | US | 2000-229358P | 20000412 | (60) |
|          |              |    |              |          |      |

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 60 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 14339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 6 OF 9 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:244853 USPATFULL TITLE: Albumin fusion proteins

INVENTOR(S): Rosen, Craig A., Laytonsville, MD, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES

Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

KIND NUMBER PATENT INFORMATION: US 2003171267 A1 20030911 US 2001-833117 A1 20010412 (9) APPLICATION INFO.:

> DATE NUMBER \_\_\_\_\_\_

PRIORITY INFORMATION:

US 2000-256931P 20001221 (60) US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT:

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 59 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 20 Drawing Page(s)

13208 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## L13 ANSWER 7 OF 9 USPATFULL on STN

тT Albumin fusion proteins

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:181414 USPATFULL TITLE: Albumin fusion proteins

Rosen, Craig A., Laytonsville, MD, UNITED STATES INVENTOR(S): Haseltine, William A., Washington, DC, UNITED STATES

NUMBER KIND DATE \_\_\_\_\_\_ US 2003125247 A1 20030703 US 2001-833041 A1 20010412 (9) PATENT INFORMATION: APPLICATION INFO.:

NUMBER DATE \_\_\_\_\_ PRIORITY INFORMATION: US 2000-256931P 20001221 (60) US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 20 Drawing Page(s)

LINE COUNT: 15235

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 8 OF 9 USPATFULL on STN

Coated particles, methods of making and using TΙ

A particle coated with a nonlamellar material such as a nonlamellar AΒ crystalline material, a nonlamellar amorphous material, or a nonlamellar semi-crystalline material includes an internal matrix core having at least one a nanostructured liquid phase, or at least on nanostructured liquid crystalline phase or a combination of the two is used for the delivery of active agents such as pharmaceuticals, nutrients, pesticides, etc. The coated particle can be fabricated by a variety of different techniques where the exterior coating is a nonlamellar material such as a nonlamellar crystalline material, a nonlamellar amorphous material, or a nonlamellar semi-crystalline material

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:159130 USPATFULL

TITLE:

Coated particles, methods of making and using

INVENTOR(S):

Anderson, David M., Colonial Heights, VA, UNITED STATES

KIND DATE NUMBER -----US 2003108743 A1 20030612 US 6638621 B2 20031028 US 2002-170237 A1 20020613 PATENT INFORMATION:

APPLICATION INFO.: 20020613 (10)

Continuation-in-part of Ser. No. US 2000-297997, filed RELATED APPLN. INFO.:

on 16 Aug 2000, GRANTED, Pat. No. US 6482517

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

WHITHAM, CURTIS & CHRISTOFFERSON, P.C., 11491 SUNSET LEGAL REPRESENTATIVE:

HILLS ROAD, SUITE 340, RESTON, VA, 20190

NUMBER OF CLAIMS: 107 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 11 Drawing Page(s)

LINE COUNT: 5538

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 9 OF 9 USPATFULL on STN L13

Multifunctional protease inhibitors and their use in treatment of TI disease

Fusion proteins of protease inhibitors are provided, in AB particular fusion proteins of alpha 1-antitrypsin (AAT) and a second protease inhibitor, such as secretory leukocyte protease inhibitor (SLPI) or tissue inhibitor of metalloproteases (TIMP). Polynucleotides encoding the fusion proteins, vectors comprising such polynucleotides, and host cells containing such vectors are also provided. Methods of making the fusion proteins of the invention are also provide, as well as methods of using the fusion proteins, for example to inhibit protease activity in a biological sample or in the treatment of an individual suffering from, or at risk for, a disease or disorder involving unwanted protease activity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:106306 USPATFULL

TITLE:

Multifunctional protease inhibitors and their use in

treatment of disease

INVENTOR (S):

Barr, Philip J., Oakland, CA, UNITED STATES Gibson, Helen, Oakland, CA, UNITED STATES

Pemberton, Philip, San Francisco, CA, UNITED STATES

NUMBER KIND DATE PATENT INFORMATION: US 2003073217 A1 20030417 A1 20011218 (10) APPLICATION INFO.: US 2001-25514

NUMBER

DATE

US 2000-256699P 20001218 (60)

US 2001-331966P 20011120 (60)

Utility DOCUMENT TYPE:

APPLICATION FILE SEGMENT:

MORRISON & FOERSTER LLP, 755 PAGE MILL RD, PALO ALTO, LEGAL REPRESENTATIVE:

CA, 94304-1018

NUMBER OF CLAIMS: 35 EXEMPLARY CLAIM: 1

PRIORITY INFORMATION:

6 Drawing Page(s) NUMBER OF DRAWINGS:

3252 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

2835 S ALBUMIN FUSION PROTEINS L1

1 S CEREBUS PROTEIN L2

0 S L1 AND L2 L3

0 S (CEREBUS PROTEIN) AND ALBUMIN L4

0 S L2 AND FUSION 1.5

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

0 S L2 L6

> FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

1 S L2 L7

8080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1 L8

5 S L8 AND L1 L9

378 S L8 AND FUSION L10

221 S L10 AND ALBUMIN L11

5 S L11 AND ALBUMIN FRAGMENT L12

9 S L11 AND SHELF-LIFE L13

=> s l11 and N-terminus fusion

0 L11 AND N-TERMINUS FUSION L14

=> s l11 and C-terminus fusion

0 L11 AND C-TERMINUS FUSION T-15

### => d l11 ti abs ibib 1-25

L11 ANSWER 1 OF 221 USPATFULL on STN

Biospecific contrast agents ΤI

Methods and apparatuses for detecting a condition of a sample (including AB cervical cancers and pre-cancers) through reflectance and/or fluorescence imaging. A sample is obtained. One or more metallic nanoparticles and/or one or more quantum dots are obtained. The one or more metallic nanoparticles and/or one or more quantum dots are coupled to one or more biomarkers of the sample that are associated with the condition. A reflectance and/or fluorescence image of the sample is then taken. The image(s) exhibit characteristic optical scattering from the one or more metallic nanoparticles and/or characteristic fluorescence excitation from the one or more quantum dots to signal the presence of the one or more biomarkers. In this way, the condition can be readily screened or diagnosed.

ACCESSION NUMBER:

2004:31276 USPATFULL

TITLE:

Biospecific contrast agents

INVENTOR(S):

Sokolov, Konstantin, Austin, TX, UNITED STATES Korgel, Brian A., Round Rock, TX, UNITED STATES Ellington, Andrew D., Austin, TX, UNITED STATES Richards-Kortum, Rebecca, Austin, TX, UNITED STATES

NUMBER KIND DATE \_\_\_\_\_\_ US 2004023415 A1 20040205 US 2003-382136 A1 20030305 (10)

NUMBER DATE \_\_\_\_\_

PRIORITY INFORMATION: DOCUMENT TYPE:

US 2002-361924P : 20020305 (60)

FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

PATENT INFORMATION: APPLICATION INFO .:

Michael C. Barrett, Esq., FULBRIGHT & JAWORSKI, L.L.P.,

600 Congress Avenue, Suite 2400, Austin, TX, 78701

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 11 Drawing Page(s)

LINE COUNT: 3948

L11 ANSWER 2 OF 221 USPATFULL on STN

Biochips for characterizing biological processes TI

This invention includes biochips for analysis of a variety of molecules, AΒ cell components and cells. Embodiments of this invention include devices and methods for the parallel and/or nearly parallel processing of biological analytes. Biochips can comprise a substrate, Raman signal-enhancing structures, and receptors selective and/or specific for the analyte(s) to be assayed. Biochips can be read using a Raman reader and can provide for rapid, sensitive, direct assays for physiological and/or pathophysiological conditions of interest.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:31155 USPATFULL

TITLE: INVENTOR(S): Biochips for characterizing biological processes Kreimer, David I., Berkeley, CA, UNITED STATES Nufert, Thomas H., Walnut Creek, CA, UNITED STATES

Ginzburg, Lev, Fremont, CA, UNITED STATES Yevin, Oleg A., Oakland, CA, UNITED STATES

NUMBER KIND DATE -----

PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.:

US 2004023293 A1 20040205 US 2002-294385 A1 20021114 (10)

Continuation-in-part of Ser. No. US 2001-925189, filed on 8 Aug 2001, PENDING Continuation-in-part of Ser. No.

US 2001-815909, filed on 23 Mar 2001, PENDING

Continuation-in-part of Ser. No. US 2000-670453, filed

on 26 Sep 2000, PENDING

NUMBER DATE 

PRIORITY INFORMATION:

US 1999-156195P 19990927 (60) US 2001-336445P 20011114 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

Sheldon R. Meyer, FLIESLER DUBB MEYER & LOVEJOY LLP, LEGAL REPRESENTATIVE:

Fourth Floor, Four Embarcadero Center, San Francisco,

CA, 94111-4156

NUMBER OF CLAIMS:

40

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

37 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 3 OF 221 USPATFULL on STN

Proteases ΤI

The invention provides human proteases (PRTS) and polynucleotides which AB identify and encode PRTS. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for diagnosing, treating, or preventing disorders associated with aberrant expression of PRTS.

ACCESSION NUMBER:

2004:31105 USPATFULL

TITLE:

Proteases

INVENTOR(S):

Henry, Yue, Sunnyvale, CA, UNITED STATES Elliott, Vicki S, San Jose, CA, UNITED STATES R Gandhi, Ameena, San Francisco, CA, UNITED STATES Lal, Preeti G, Santa Clara, CA, UNITED STATES Au-Young, Janice, Brisbane, CA, UNITED STATES Tribouley, Catherine M, San Francisco, CA, UNITED

STATES Delegeane, Angelo M, Milpitas, CA, UNITED STATES Baughn, Mariah R, San Leandro, CA, UNITED STATES Nguyen, Danniel B, San Jose, CA, UNITED STATES Lee, Ernestine A, Albany, CA, UNITED STATES Hafalia, April J A, Daly City, CA, UNITED STATES Khan, Farrah A, Des Plaines, IL, UNITED STATES Chawla, Narinder K, Union City, CA, UNITED STATES Yao, Monique G, Carmel, IN, UNITED STATES Lu, Dyung Aina M, San Jose, CA, UNITED STATES Arvizu, Chandra S, San Jose, CA, UNITED STATES Tang, Y Tom, San Jose, CA, UNITED STATES Walsh, Roderick T, Canterbury, UNITED KINGDOM Azimzai, Yalda, Oakland, CA, UNITED STATES Lu, Yan, Palo Alto, CA, UNITED STATES Ramkumar, Jayalaxmi, Fremont, CA, UNITED STATES Xu, Yuming, Mountain View, CA, UNITED STATES Reddy, Roopa, Sunnyvale, CA, UNITED STATES Das, Debopriya, Mountain View, CA, UNITED STATES Kearney, Liam, San Francisco, CA, UNITED STATES Kallick, Deborah A, Galveston, TX, UNITED STATES

|                     | NUMBER          | KIND | DATE     |      |
|---------------------|-----------------|------|----------|------|
|                     |                 |      |          |      |
| PATENT INFORMATION: | US 2004023243   | A1   | 20040205 |      |
| APPLICATION INFO.:  | US 2003-311035  | A1   | 20030519 | (10) |
|                     | WO 2001-US19178 |      | 20010613 |      |
| DOCUMENT TYPE:      | Utility         |      |          |      |
| PILE CECMENT.       | A DDI.TCATTON   |      |          |      |

LEGAL REPRESENTATIVE:

INCYTE CORPORATION (formerly known as Incyte, Genomics,

Inc.), 3160 PORTER DRIVE, PALO ALTO, CA, 94304

NUMBER OF CLAIMS: 116 EXEMPLARY CLAIM: LINE COUNT: 8891

L11 ANSWER 4 OF 221 USPATFULL on STN

Novel human gene relating to respiratory diseases, obesity, and ΤI inflammatory bowel disease

This invention relates to genes identified from human chromosome ΔR 20p13-p12, which are associated with various diseases, including asthma. The invention also relates to the nucleotide sequences of these genes, isolated nucleic acids comprising these nucleotide sequences, and isolated polypeptides or peptides encoded thereby. The invention further relates to vectors and host cells comprising the disclosed nucleotide sequences, or fragments thereof, as well as antibodies that bind to the encoded polypeptides or peptides. Also related are ligands that modulate

the activity of the disclosed genes or gene products. In addition, the invention relates to methods and compositions employing the disclosed nucleic acids, polypeptides or peptides, antibodies, and/or ligands for use in diagnostics and therapeutics for asthma and other diseases.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

INVENTOR(S):

2004:31077 USPATFULL

TITLE:

Novel human gene relating to respiratory diseases,

obesity, and inflammatory bowel disease

Keith, Tim, Bedford, MA, UNITED STATES

Little, Randall D., Newtonville, MA, UNITED STATES

Eerdewegh, Paul Van, Weston, MA, UNITED STATES

Dupuis, Josee, Newton, MA, UNITED STATES

Del Mastro, Richard G., Norfolk, MA, UNITED STATES

Simon, Jason, Westfield, NJ, UNITED STATES Allen, Kristina, Hopkinton, MA, UNITED STATES Pandit, Sunil, Gaithersburg, MD, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_\_

PATENT INFORMATION: APPLICATION INFO.:

US 2004023215 A1 20040205 US 2002-126022 A1 20020419 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2001-834597, filed

on 13 Apr 2001, PENDING Continuation-in-part of Ser. No. US 2000-548797, filed on 13 Apr 2000, PENDING

DATE NUMBER -------

PRIORITY INFORMATION: US 1999-129391P 19990413 (60)

DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

MORGAN & FINNEGAN, L.L.P., 345 Park Avenue, New York,

NY, 10154-0053

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

157 Drawing Page(s)

LINE COUNT:

20001

73

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 5 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI AB

The present invention relates to novel proteins. More specifically, isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

ACCESSION NUMBER:

2004:25127 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES

KIND DATE NUMBER -----US 2004018969 A1 20040129

PATENT INFORMATION:

| APPLICATION INFO.:    | US 2001-764875  | A1 20010117   | (9) |
|-----------------------|---|---|-----|
|                       | NUMBER  | DATE.   |     |
| PRIORITY INFORMATION: | US 2000-179065P<br>US 2000-180628P<br>US 2000-214886P<br>US 2000-217487P<br>US 2000-225758P<br>US 2000-220963P<br>US 2000-217496P<br>US 2000-217496P<br>US 2000-225447P<br>US 2000-225757P<br>US 2000-226868P | 20000131 (60)<br>20000204 (60)<br>20000628 (60)<br>20000711 (60)<br>20000726 (60)<br>20000711 (60)<br>20000714 (60)<br>20000814 (60)<br>20000814 (60)<br>20000814 (60)<br>20000814 (60) |     |
|                       | US 2000-216647P US 2000-225267P US 2000-216880P US 2000-225270P US 2000-251869P US 2000-235834P US 2000-234274P US 2000-234223P US 2000-228924P US 2000-224518P US 2000-236369P                               | 20000707 (60)<br>20000814 (60)<br>20000707 (60)<br>20000814 (60)<br>20001208 (60)<br>20000927 (60)<br>20000921 (60)<br>20000921 (60)<br>20000830 (60)<br>20000814 (60)<br>20000929 (60) |     |
|                       | US 2000-224519P<br>US 2000-220964P<br>US 2000-241809P<br>US 2000-249299P<br>US 2000-236327P<br>US 2000-241785P<br>US 2000-244617P<br>US 2000-225268P<br>US 2000-236368P<br>US 2000-251856P                    | 20000814 (60)<br>20000726 (60)<br>20001020 (60)<br>20001117 (60)<br>20000929 (60)<br>20001020 (60)<br>20001101 (60)<br>20000814 (60)<br>20000929 (60)<br>20001208 (60)                  |     |
|                       | US 2000-251836P<br>US 2000-251868P<br>US 2000-229344P<br>US 2000-234997P<br>US 2000-229345P<br>US 2000-229345P<br>US 2000-229513P<br>US 2000-231413P<br>US 2000-231413P<br>US 2000-236367P                    | 20001208 (60)<br>20000901 (60)<br>20000925 (60)   |     |
|                       | US 2000-237039P<br>US 2000-237038P<br>US 2000-236370P<br>US 2000-236802P<br>US 2000-237037P<br>US 2000-237040P<br>US 2000-240960P<br>US 2000-239935P<br>US 2000-239937P<br>US 2000-241787P                    | 20001002 (60)<br>20001002 (60)<br>20000929 (60)<br>20001002 (60)<br>20001002 (60)<br>20001002 (60)<br>20001013 (60)<br>20001013 (60)<br>20001020 (60)                                   |     |
|                       | US 2000-246474P<br>US 2000-246532P<br>US 2000-249216P<br>US 2000-249210P<br>US 2000-226681P<br>US 2000-225759P<br>US 2000-225213P<br>US 2000-227182P  | 20001108 (60)<br>20001108 (60)<br>20001117 (60)<br>20001117 (60)<br>20000822 (60)<br>20000814 (60)<br>20000814 (60)<br>20000822 (60)  |     |

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US 2000-225214P
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                    20001108
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DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

24 1

LINE COUNT:

38235

ANSWER 6 OF 221 USPATFULL on STN L11

TT Molecules for diagnostics and therapeutics

The present invention provides purified human polynucleotides for AB diagnostics and therapeutics (dithp). Also encompassed are the polypeptides (DITHP) encoded by dithp. The invention also provides for the use of dithp, or complements, oligonucleotides, or fragments thereof in diagnostic assays. The invention further provides for vectors and host cells containing dithp for the expression of DITHP. The invention additionally provides for the use of isolated and purified DITHP to induce antibodies and to screen libraries of compounds and the use of anti-DITHP antibodies in diagnostic assays. Also provided are microarrays containing dithp and methods of use.

ACCESSION NUMBER:

2004:18785 USPATFULL

TITLE: INVENTOR (S): Molecules for diagnostics and therapeutics Hodgson, David M., Ann Arbor, MI, UNITED STATES Lincoln, Stephen E., Potomac, MD, UNITED STATES Russo, Frank D., Sunnyvale, CA, UNITED STATES Albany, Peter A., Berkeley, CA, UNITED STATES Banville, Steve C., Sunnyvale, CA, UNITED STATES Bratcher, Shawn R., Mountain View, CA, UNITED STATES Dufour, Gerard E., Castro Valley, CA, UNITED STATES Cohen, Howard J., Palo Alto, CA, UNITED STATES Rosen, Bruce H., Menlo Park, CA, UNITED STATES

Chalup, Michael S., Livingston, TX, UNITED STATES Jackson, Jennifer L., Santa Cruz, CA, UNITED STATES Jones, Anissa L., San Jose, CA, UNITED STATES

Yu, Jimmy Y., Fremont, CA, UNITED STATES

Greenawalt, Lila B., San Jose, CA, UNITED STATES Panzer, Scott R., Sunnyvale, CA, UNITED STATES Roseberry Lincoln, Ann M., Potomac, MD, UNITED STATES

Wright, Rachel J., Merivale, NEW ZEALAND

Daniels, Susan E., Mountain View, CA, UNITED STATES Incyte Corporation, Palo Alto, CA, UNITED STATES (U.S.

corporation)

PATENT ASSIGNEE(S):

|      | NUMBER      | KIND | DATE     |  |
|------|-------------|------|----------|--|
|      |             |      |          |  |
| US   | 2004014087  | A1   | 20040122 |  |
| יזני | 2002 270020 | 7.1  | 2002022  |  |

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

(10) . US 2003-378029 **A1** 20030228 Continuation-in-part of Ser. No. US 2001-980285, filed on 30 Nov 2001, PENDING A 371 of International Ser. No. WO 2000-US15404, filed on 31 May 2000, PENDING

DATE NUMBER \_\_\_\_\_\_ 19990805 (60) US 1999-147500P PRIORITY INFORMATION: 19990805 (60) US 1999-147542P 19990805 (60) US 1999-147541P 19990805 (60) US 1999-147824P 19990805 (60) US 1999-147547P 19990805 (60) US 1999-147530P 19990805 (60) US 1999-147536P 19990805 (60) US 1999-147520P 19990805 (60) US 1999-147527P 19990805 (60) US 1999-147549P 19990804 (60) US 1999-147377P 19990804 (60) US 1999-147436P US 1999-137411P 19990603 (60) US 1999-137396P 19990603 (60) US 1999-137417P 19990603 (60) 19990603 (60) US 1999-137337P 19990602 (60) US 1999-137173P 19990602 (60) US 1999-137114P US 1999-137259P 19990602 (60) 19990602 (60) US 1999-137113P 19990602 (60) US 1999-137260P 19990602 (60) US 1999-137258P 19990602 (60) US 1999-137109P 19990601 (60) US 1999-137161P Utility

DOCUMENT TYPE: . FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

INCYTE CORPORATION (formerly known as Incyte, Genomics,

Inc.), 3160 PORTER DRIVE, PALO ALTO, CA, 94304

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 14819

L11 ANSWER 7 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI

The present invention relates to novel proteins. More specifically, AB isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

ACCESSION NUMBER:

2004:18737 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

PATENT ASSIGNEE(S): STATES, 20850 (U.S. corporation)

| KIND | DATE     |
|------|----------|
|      |          |
| A1   | 20040122 |
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PATENT INFORMATION: APPLICATION INFO.:

US 2002-158057 A1 20020531 (10)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2001-764890, filed on 17

Jan 2001, PENDING

|                       | NUMBER                             | DATE                           |
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DOCUMENT TYPE: FILE SEGMENT:

LINE COUNT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

1 26776

L11 ANSWER 8 OF 221 USPATFULL on STN

TI Albumin fusion proteins

AB The present invention encompasses albumin fusion

proteins. Nucleic acid molecules encoding the **albumin fusion** proteins of the invention are also encompassed by the
invention, as are vectors containing these nucleic acids, host cells
transformed with these nucleic acids vectors, and methods of making the **albumin fusion** proteins of the invention and using

these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating,

preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL Albumin fusion proteins

TITLE: INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2004010134  | A1   | 20040115 |     |
| APPLICATION INFO.:  | US 2001-833245 | A1   | 20010412 | (9) |

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 9 OF 221 USPATFULL on STN

TI 7 Human ovarian and ovarian cancer associated proteins

AB This invention relates to newly identified ovarian or ovarian cancer related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "ovarian cancer antigens",

and the use of such ovarian antigens for detecting disorders of the reproductive system, particularly the presence of ovarian cancer and ovarian cancer metastases. This invention relates to ovarian cancer antigens as well as vectors, host cells, antibodies directed to ovarian cancer antigens and the recombinant methods and synthetic methods for producing the same. Also provided are diagnostic methods for detecting, treating, preventing and/or prognosing disorders related to the ovary, including ovarian cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of ovarian cancer antigens of the invention. The present invention further relates to inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13598 USPATFULL

AΒ

7 Human ovarian and ovarian cancer associated proteins

INVENTOR(S):

Birse, Charles E., North Potomac, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES

| NUMBER  | KIND     | DATE                             |      |
|---|----------|----------------------------------|------|
| US 2004010121<br>US 2003-333900<br>WO 2001-US8585 | A1<br>A1 | 20040115<br>20030124<br>20010316 | (10) |

APPLICATION INFO.: DOCUMENT TYPE:

PATENT INFORMATION:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

23 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 16023 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 10 OF 221 USPATFULL on STN

Use of bioactive glass compositions to stimulate osteoblast production ΤI Compositions comprising bioactive glass compositions or extracts thereof which include ions in an appropriate concentration and ratio that they enhance osteoblast production, and methods of preparation and use thereof, are disclosed. The compositions can be included in implantable devices that are capable of inducing tissue formation in autogeneic, allogeneic and xenogeneic implants, for example as coatings and/or matrix materials. Examples of such devices include prosthetic implants, sutures, stents, screws, plates, tubes, and the like. Aqueous extracts of the bioactive glass compositions, which extracts are capable of stimulating osteoblast production, are also disclosed. The compositions can be used, for example, to induce local tissue formation from a progenitor cell in a mammal, for accelerating allograft repair in a mammal, for promoting in vivo integration of an implantable prosthetic device to enhance the bond strength between the prosthesis and the existing target tissue at the joining site, and for treating tissue degenerative conditions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13078 USPATFULL

TITLE:

Use of bioactive glass compositions to stimulate

osteoblast production

INVENTOR(S):

Hench, Larry L, London, UNITED KINGDOM Polak, Julia M, London, UNITED KINGDOM Buttery, Lee D.k., London, UNITED KINGDOM

Xynos, Ioannis D, Nafplion, GREECE

Maroothynaden, Jason, London, UNITED KINGDOM

KIND DATE NUMBER

20040115 PATENT INFORMATION: US 2004009598 A1 A1 20030707 (10)

US 2003-332731 APPLICATION INFO.:

20010711 WO 2001-US21801

Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT:

BURNS DOANE SWECKER & MATHIS L L P, POST OFFICE BOX LEGAL REPRESENTATIVE:

1404, ALEXANDRIA, VA, 22313-1404

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 1301 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 11 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ΤI

The present invention relates to novel polynucleotides associated with AB the plasma membrane, the polypeptides encoded by these polynucleotides herein collectively referred to as "plasma membrane associated antigens," and antibodies that immunospecifically bind these polypeptides, and the use of such plasma membrane associated polynucleotides, antigens, and antibodies for detecting, treating, preventing and/or prognosing disorders related to these novel polypeptides. More specifically, isolated nucleic acid molecules are provided encoding novel plasma membrane associated polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing these plasma membrane associated polynucleotides, polypeptides, and/or antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the novel polypeptides of the invention. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The invention further relates to methods and/or compositions for inhibiting or promoting the production and/or function of the polypeptides of the invention.

2004:12971 USPATFULL ACCESSION NUMBER:

Nucleic acids, proteins, and antibodies TITLE:

Birse, Charles E., North Potomac, MD, UNITED STATES INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_

US 2004009491 A1 20040115 US 2002-264237 A1 20021004 PATENT INFORMATION: (10) APPLICATION INFO .:

Continuation-in-part of Ser. No. WO 2001-US16450, filed RELATED APPLN. INFO.:

on 18 May 2001, PENDING

DATE NUMBER \_\_\_\_\_

US 2000-205515P 20000519 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1 LINE COUNT: 18144

L11 ANSWER 12 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI

The present invention relates to novel musculoskeletal system related AB polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "musculoskeletal system antigens," and the use of such musculoskeletal system antigens for detecting disorders of

the musculoskeletal system, particularly the presence of cancer and cancer metastases. More specifically, isolated musculoskeletal system associated nucleic acid molecules are provided encoding novel musculoskeletal system associated polypeptides. Novel musculoskeletal system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human musculoskeletal system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the musculoskeletal system, including cancer of musculoskeletal tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:12968 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_\_

PATENT INFORMATION: APPLICATION INFO.:

PATENT ASSIGNEE(S):

US 2004009488 A1 20040115 US 2002-242515 A1 20020913 (10)

DATE

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2001-764877, filed on 17

Jan 2001, PENDING

NUMBER

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Utility
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DOCUMENT TYPE:

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: LINE COUNT: 32038

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### ANSWER 13 OF 221 USPATFULL on STN L11

TΙ Methods for the treatment of carcinoma

The invention concerns compositions and methods for the diagnosis and AB treatment of neoplastic cell growth and proliferation in mammals, including humans. The invention is based upon the identification of genes that are amplified in the genome of tumor cells, such as renal cell carcinoma. Such gene amplification is expected to be associated with the overexpression of the gene product as compared to normal cells of the same tissue type and contribute to tumorigenesis. Accordingly, the proteins encoded by the amplified genes are believed to be useful targets for the diagnosis and/or treatment (including prevention) of certain cancers, such as renal cell carcinoma, and may act as predictors of the prognosis of tumor treatment. The present invention is directed to novel methods of diagnosing and treating tumor, such as renal cell carcinoma or Wilms tumor.

ACCESSION NUMBER:

2004:12653 USPATFULL

TITLE:

Methods for the treatment of carcinoma

INVENTOR(S):

Gerritsen, Mary E., San Mateo, CA, UNITED STATES

Peale, Franklin V., JR., San Carlos, CA, UNITED STATES

Wu, Thomas D., San Francisco, CA, UNITED STATES

PATENT ASSIGNEE(S):

GENENTECH, INC. (U.S. corporation)

NUMBER KIND \_\_\_\_\_\_ A1 20040115 A1 20030221 US 2004009171 PATENT INFORMATION:

APPLICATION INFO .:

US 2003-372683 20030221 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2002-271690, filed

on 16 Oct 2002, PENDING

NUMBER DATE

PRIORITY INFORMATION:

US 2001-344534P 20011018 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

GENENTECH, INC., 1 DNA WAY, SOUTH SAN FRANCISCO, CA,

94080

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT:

AΒ

57 1 6662

L11 ANSWER 14 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ΤI

The present invention relates to novel ovarian related polynucleotides, the polypeptides encoded by these polynucleotides herein collectively referred to as "ovarian antigens," and antibodies that immunospecifically bind these polypeptides, and the use of such ovarian polynucleotides, antigens, and antibodies for detecting, treating, preventing and/or prognosing disorders of the reproductive system, particularly disorders of the ovaries and/or breast, including, but not limited to, the presence of ovarian and/or breast cancer and ovarian and/or breast cancer metastases. More specifically, isolated ovarian nucleic acid molecules are provided encoding novel ovarian polypeptides. Novel ovarian polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human ovarian polynucleotides, polypeptides, and/or antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the ovaries and/or breast, including ovarian and/or breast cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The invention further relates to methods and/or compositions for inhibiting or promoting the production and/or function of the polypeptides of the invention.

ACCESSION NUMBER:

2004:7345 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

KIND

INVENTOR(S):

Birse, Charles E., North Potomac, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES

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|---------------------|----------------|-------|-----------|---|
|                     |                |       | <b></b> - |   |
| PATENT INFORMATION: | US 2004005579  | A1    | 20040108  |   |
| APPLICATION INFO.:  | US 2002-264049 | A1    | 20021004  | ( |

MIMBER

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. WO 2001-US18569, filed

DATE

#### on 7 Jun 2001, PENDING

DATE NUMBER \_\_\_\_\_\_ US 2000-209467P 20000607 (60)

PRIORITY INFORMATION: Utility DOCUMENT TYPE:

APPLICATION FILE SEGMENT:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 18130 LINE COUNT:

AB

L11 ANSWER 15 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ΤI

The present invention relates to novel proteins. More specifically, isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2004:7343 USPATFULL ACCESSION NUMBER:

PATENT INFORMATION:

Nucleic acids, proteins, and antibodies

Rosen, Craig A., Laytonsville, MD, UNITED STATES INVENTOR (S):

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES

20001208 (60)

20000927 (60)

Human Genome Sciences, Inc., Rockville, MD, UNITED PATENT ASSIGNEE(S):

STATES (U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_ US 2004005577 A1 20040108 US 2002-242747 A1 20020913 (10)

APPLICATION INFO .: Continuation of Ser. No. US 2001-764881, filed on 17

RELATED APPLN. INFO.:

Jan 2001, PENDING

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Utility
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DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1 LINE COUNT: 27694

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 16 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies

The present invention relates to novel cardiovascular system related AB polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "cardiovascular system antigens," and the use of such cardiovascular system antigens for detecting disorders of the cardiovascular system, particularly the presence of cancer of cardiovascular system tissues and cancer metastases. More specifically, isolated cardiovascular system associated nucleic acid molecules are provided encoding novel cardiovascular system associated polypeptides. Novel cardiovascular system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human cardiovascular system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the cardiovascular system, including cancer of cardiovascular system tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:7341 USPATFULL

TITLE:

ΤI

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

PATENT ASSIGNEE(S):

KIND DATE NUMBER \_\_\_\_\_\_

PATENT INFORMATION: APPLICATION INFO.:

US 2004005575 20040108 A1 A1 US 2002-227577 20020826

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2002-91504, filed on 7 Mar 2002, PENDING Continuation of Ser. No. US 2001-764869,

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filed on 17 Jan 2001, ABANDONED

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US 2000-205515P
                    20010105 (60)
US 2001-259678P
Utility
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DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24
EXEMPLARY CLAIM: 1
LINE COUNT: 28742

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 17 OF 221 USPATFULL on STN

TI Functional MRI agents for cancer imaging

AB The invention relates to novel magnetic resonance imaging contrast agents for imaging cancer.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2004:4285 USPATFULL ACCESSION NUMBER:

Functional MRI agents for cancer imaging TITLE:

Meade, Thomas J., Altadena, CA, United States INVENTOR(S): Fraser, Scott, La Canada, CA, United States

Jacobs, Russell, Arcadia, CA, United States

Research Corporation Technologies, Inc., Tucson, AZ, PATENT ASSIGNEE(S):

United States (U.S. corporation)

KIND DATE NUMBER \_\_\_\_\_\_

PATENT INFORMATION:

B1 20040106 US 6673333 20001117 (9)

US 2000-715859 APPLICATION INFO.:

> DATE NUMBER \_\_\_\_\_\_

US 2000-201816P 20000504 (60) PRIORITY INFORMATION:

Utility DOCUMENT TYPE: GRANTED FILE SEGMENT:

Hartley, Michael G. PRIMARY EXAMINER:

Dorsey & Whitney LLP, Silva, Robin M., Kossiak, Renee LEGAL REPRESENTATIVE:

NUMBER OF CLAIMS: 10 EXEMPLARY CLAIM:

7 Drawing Figure(s); 5 Drawing Page(s) NUMBER OF DRAWINGS:

2422 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 18 OF 221 USPATFULL on STN

TI 50 human secreted proteins

The present invention relates to novel human secreted proteins and AB isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating diseases, disorders, and/or conditions related to these novel human secreted proteins.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:2568 USPATFULL

TITLE:

50 human secreted proteins

Moore, Paul A., Germantown, MD, UNITED STATES INVENTOR(S):

Ruben, Steven M., Olney, MD, UNITED STATES LaFleur, David W., Washington, DC, UNITED STATES Shi, Yanggu, Gaithersburg, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES Olsen, Henrik S., Gaithersburg, MD, UNITED STATES Ebner, Reinhard, Gaithersburg, MD, UNITED STATES Brewer, Laurie A., St. Paul, MN, UNITED STATES

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD (U.S.

corporation)

KIND NUMBER DATE \_\_\_\_\_\_

PATENT INFORMATION:

US 2004002591 A1 20040101 US 2002-47021 A1 20020117 (10)

APPLICATION INFO .:

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2000-722329, filed

on 28 Nov 2000, PENDING Continuation of Ser. No. US

1999-262109, filed on 4 Mar 1999, ABANDONED

Continuation-in-part of Ser. No. WO 1998-US18360, filed

on 3 Sep 1998, PENDING

NUMBER DATE PRIORITY INFORMATION:

US 2001-262066P 20010118 (60) 19970905 (60) US 1997-57626P

19970905 (60) US 1997-57663P

US 1997-57669P 19970905 (60)

19970912 (60) US 1997-58666P

19970912 (60) US 1997-58667P

19970912 (60) US 1997-58973P

US 1997-58974P 19970912 (60) 19980622 (60) US 1998-90112P

Utility DOCUMENT TYPE:

FILE SEGMENT: APPLICATION

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 23

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Page(s)

LINE COUNT: 33379

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 19 OF 221 USPATFULL on STN

Novel human gene relating to respiratory diseases, obesity, and TI

inflammatory bowel disease

This invention relates to genes identified from human chromosome AΒ 20p13-p12, which are associated with various diseases, including asthma.

The invention also relates to the nucleotide sequences of these genes, isolated nucleic acids comprising these nucleotide sequences, and isolated polypeptides or peptides encoded thereby. The invention further relates to vectors and host cells comprising the disclosed nucleotide sequences, or fragments thereof, as well as antibodies that bind to the encoded polypeptides or peptides. Also related are ligands that modulate the activity of the disclosed genes or gene products. In addition, the invention relates to methods and compositions employing the disclosed nucleic acids, polypeptides or peptides, antibodies, and/or ligands for use in diagnostics and therapeutics for asthma and other diseases.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2004:2447 USPATFULL ACCESSION NUMBER:

Novel human gene relating to respiratory diseases, TITLE:

obesity, and inflammatory bowel disease

Keith, Tim, Bedford, MA, UNITED STATES INVENTOR(S):

Little, Randall D., Newtonville, MA, UNITED STATES

Eerdewegh, Paul Van, Weston, MA, UNITED STATES

Dupuis, Josee, Newton, MA, UNITED STATES

Del Mastro, Richard G., Norfolk, MA, UNITED STATES

Simon, Jason, Westfield, NJ, UNITED STATES Allen, Kristin, Hopkinton, MA, UNITED STATES Pandit, Sunil, Gaithersburg, MD, UNITED STATES

| NUMBER KIND | DATE |
|-------------|------|
|-------------|------|

PATENT INFORMATION: US 2004002470 A1 20040101

A1 20021017 (10) APPLICATION INFO.: US 2002-277216

Continuation-in-part of Ser. No. US 2002-126022, filed RELATED APPLN. INFO.: on 19 Apr 2002, PENDING Continuation-in-part of Ser. No. US 2001-834597, filed on 13 Apr 2001, PENDING

Continuation-in-part of Ser. No. US 2000-548797, filed

on 13 Apr 2000, PENDING

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

MORGAN & FINNEGAN, L.L.P., 345 PARK AVENUE, NEW YORK, LEGAL REPRESENTATIVE:

NY, 10154

NUMBER OF CLAIMS: 45 EXEMPLARY CLAIM: 1

162 Drawing Page(s) NUMBER OF DRAWINGS:

15810 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 20 OF 221 USPATFULL on STN

Detection and modulation of Slit and roundabount (Robo) mediated TI

angiogenesis and uses thereof

This invention is generally in the field of methods for diagnosis, AB treatment and prevention of various disorders involving the Slit2 mediated angiogenesis.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:335332 USPATFULL

TITLE:

Detection and modulation of Slit and roundabount (Robo)

mediated angiogenesis and uses thereof

INVENTOR (S):

Geng, Jian-Guo, Portage, MI, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION:

US 2003236210 A1 20031225

APPLICATION INFO.:

US 2003-386386 A1 20030310 (10)

NUMBER DATE -----

PRIORITY INFORMATION:

US 2002-362485P 20020308 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Peng Chen, Morrison & Foerster LLP, Suite 500, 3811

Valley Centre Drive, San Diego, CA, 92130-2332

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

4 Drawing Page(s)

NUMBER OF DRAWINGS: LINE COUNT:

AΒ

1337

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 21 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI

The present invention relates to novel excretory system related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "excretory system antigens," and the use of such excretory system antigens for detecting disorders of the excretory system, particularly the presence of cancer of excretory system tissues and cancer metastases. More specifically, isolated excretory system associated nucleic acid molecules are provided encoding novel excretory system associated polypeptides. Novel excretory system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human excretory system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the excretory system, including cancer of excretory system tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2003:334955 USPATFULL ACCESSION NUMBER:

TITLE: Nucleic acids, proteins, and antibodies

Rosen, Craig A., Laytonsville, MD, UNITED STATES INVENTOR (S):

Ruben, Steven M., Olney, MD, UNITED STATES Barash, Steven C., Rockville, MD, UNITED STATES PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD, 20850 (U.S. corporation)

| NUMBER                           | KIND    | DATE     |      | ,     |       |
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| US 2003235831<br>US 2002-242355  |         | 20031225 | (10) |       |       |
| Continuation of Jan 2001, PENDIN | Ser. No |          |      | filed | on 17 |

APPLICATION INFO.: RELATED APPLN. INFO.:

PATENT INFORMATION:

PRIORITY INFORMATION:

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DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT:

22457

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 22 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI

The present invention relates to novel proteins. More specifically, AB isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:334953 USPATFULL

TITLE: INVENTOR (S): Nucleic acids, proteins, and antibodies Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES Birse, Charles E., North Potomac, MD, UNITED STATES

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES (U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_\_ US 2003235829 20031225 PATENT INFORMATION: A1 20020826 APPLICATION INFO.: US 2002-227646 **A1** 

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2001-860670, filed on 21 May 2001, PENDING Continuation-in-part of Ser. No. WO

2001-US1346, filed on 17 Jan 2001, PENDING

NUMBER DATE

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Utility
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DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24
EXEMPLARY CLAIM: 1
LINE COUNT: 20415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 23 OF 221 USPATFULL on STN

TI Compositions and methods for systemic inhibition of cartilage degradation

AB Methods and compositions for inhibiting articular cartilage degradation. The compositions preferably include multiple chondroprotective agents, including at least one agent that promotes cartilage anabolic activity and at least one agent that inhibits cartilage catabolism. The compositions may also include one or more pain and inflammation

inhibitory agents. The compositions may be administered systemically, such as to treat patients at risk of cartilage degradation at multiple joints, and suitably may be formulated in a carrier or delivery vehicle that is targeted to the joints. Alternatively the compositions may be injected or infused directly into the joint.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:334713 USPATFULL

TITLE:

Compositions and methods for systemic inhibition of

cartilage degradation

INVENTOR (S):

Demopulos, Gregory A., Mercer Island, WA, UNITED STATES Palmer, Pamela Pierce, San Francisco, CA, UNITED STATES

Herz, Jeffrey M., Mill Creek, WA, UNITED STATES

PATENT ASSIGNEE(S):

Omeros Corporation (U.S. corporation)

| NUMBER | KIND | DATE |
|--------|------|------|
|        |      |      |

PATENT INFORMATION:

US 2003235589 A1 20031225

APPLICATION INFO .: RELATED APPLN. INFO.: US 2003-356649 A1 20030131 (10) Continuation-in-part of Ser. No. US 2002-31546, filed

on 18 Jan 2002, PENDING A 371 of International Ser. No.

WO 2000-US19864, filed on 21 Jul 2000, PENDING

Continuation-in-part of Ser. No. US 2001-839633, filed on 20 Apr 2001, PENDING Continuation-in-part of Ser. No. WO 1999-US26330, filed on 5 Nov 1999, PENDING Continuation-in-part of Ser. No. WO 1999-US24625, filed

on 20 Oct 1999, PENDING

| NUMBER | DATE |
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PRIORITY INFORMATION:

US 2002-353552P 20020201 (60) US 1999-144904P 19990721 (60) US 1998-107256P 19981105 (60) 19981020 (60) US 1998-105026P

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

OMEROS MEDICAL SYSTEMS, INC., 1420 FIFTH AVENUE, SUITE

2675, SEATTLE, WA, 98101

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

155

NUMBER OF DRAWINGS:

9 Drawing Page(s)

LINE COUNT:

6575

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## ANSWER 24 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TI

The present invention relates to novel endocrine related polynucleotides AΒ and the polypeptides encoded by these polynucleotides herein collectively known as "endocrine antigens," and the use of such endocrine antigens for detecting disorders of the endocrine system, particularly the presence of cancers of the endocrine system and endocrine cancer metastases. More specifically, isolated endocrine associated nucleic acid molecules are provided encoding novel endocrine associated polypeptides. Novel endocrine polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human endocrine associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the endocrine system, including cancers of the endocrine system, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the

production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:330759 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Ruben, Steven M., Olney, MD, UNITED STATES Barash, Steven C., Rockville, MD, UNITED STATES

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD (U.S.

corporation)

NUMBER KIND DATE \_\_\_\_\_ US 2003232975 A1 A1

PATENT INFORMATION: APPLICATION INFO.:

20031218 20020214

RELATED APPLN. INFO.:

(10) Continuation of Ser. No. US 2001-764895, filed on 17

Jan 2001, ABANDONED мити

|          |              |    | NUMBER                       |   | DATE              |              |
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US 2001-259678P
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DOCUMENT TYPE:

Utility

FILE SEGMENT: LEGAL REPRESENTATIVE: APPLICATION HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

24

21828

EXEMPLARY CLAIM: LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 25 OF 221 USPATFULL on STN

TT Proteases

The invention provides human proteases (PRTS) and polynucleotides which AΒ identify and encode PRTS. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for diagnosing, treating, or preventing disorders associated with aberrant expression of PRTS.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:330138 USPATFULL

TITLE:

Proteases

INVENTOR(S):

Delegeane, Angelo M., Milpitas, CA, UNITED STATES Gandhi, Ameena R., San Francisco, CA, UNITED STATES Hafalia, April J. A., Santa Clara, CA, UNITED STATES Lu, Dyung Aina M., San Jose, CA, UNITED STATES Arvizu, Chandra S., San Jose, CA, UNITED STATES Tribouley, Catherine M., San Francisco, CA, UNITED STATES

Das, Debopriya, Mountain View, CA, UNITED STATES Kallick, Deborah A., Portola Valley, CA, UNITED STATES Nguyen, Danniel B., San Jose, CA, UNITED STATES Lee, Ernestine A., Castro Valley, CA, UNITED STATES Khan, Farrah A., Glen View, IL, UNITED STATES Yue, Henry, Sunnyvale, CA, UNITED STATES Au-Young, Janice, Brisbane, CA, UNITED STATES Griffin, Jennifer A., Fremont, CA, UNITED STATES

Policky, Jennifer L., San Jose, CA, UNITED STATES

Ramkumar, Jayalaxmi, Fremont, CA, UNITED STATES Yang, Junming, San Jose, CA, UNITED STATES Thangavelu, Kavitha, Mountain View, CA, UNITED STATES Ding, Li, Creve Coeur, MO, UNITED STATES Kearney, Liam, San Francisco, CA, UNITED STATES Baughn, Mariah R., San Leandro, CA, UNITED STATES Borowsky, Mark L., Redwood City, CA, UNITED STATES Sanjanwala, Madhusudan, Los Altos, CA, UNITED STATES Yao, Monique G., Carmel, IN, UNITED STATES Burford, Neil, Durham, CT, UNITED STATES Chawla, Narinder K., Union City, CA, UNITED STATES Lal, Preeti G., Santa Clara, CA, UNITED STATES Lee, Sally, San Jose, CA, UNITED STATES Todd, Stephen, San Francisco, CA, UNITED STATES Lo, Terence P., Foster City, CA, UNITED STATES Tang, Y. Tom, San Jose, CA, UNITED STATES Elliott, Vicki S., San Jose, CA, UNITED STATES Azimzai, Yalda, Oakland, CA, UNITED STATES Lu, Yan, Palo Alto, CA, UNITED STATES

PATENT ASSIGNEE(S):

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.: Incyte Genomics, Inc., Palo Alto, CA (U.S. corporation)

| NUMBER           | KIND       | DATE       |          |       |    |    |
|------------------|------------|------------|----------|-------|----|----|
|                  |            |            |          |       |    |    |
| US 2003232349    | A1         | 20031218   |          |       |    |    |
| US 2002-274639   | A1         | 20021018   | (10)     |       |    |    |
| Continuation of  | Ser. No.   | . WO 2001- | US22397, | filed | on | 17 |
| Jul 2001, PENDIN | I <b>G</b> |            |          |       |    |    |

|          | ·            |     | NUMBER       | DATE     |      |
|----------|--------------|-----|--------------|----------|------|
|          |              |     |              |          |      |
| PRIORITY | INFORMATION: | US  | 2000-220063P | 20000721 | (60) |
|          |              | US  | 2000-221680P | 20000728 | (60) |
|          |              | US  | 2000-223544P | 20000804 | (60) |
|          |              | US  | 2000-224717P | 20000811 | (60) |
|          |              | US  | 2000-225988P | 20000816 | (60) |
|          |              | US  | 2000-227568P | 20000823 | (60) |
| DOCUMENT | TYPE:        | Uti | lity         | •        |      |

FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: INCYTE CORPORATION (formerly known as Incyte, Genomics,

Inc.), 3160 PORTER DRIVE, PALO ALTO, CA, 94304

NUMBER OF CLAIMS: 86
EXEMPLARY CLAIM: 1
LINE COUNT: 8959

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

NEWS PHONE

NEWS WWW

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\* STN Columbus

FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004

=> file medline, uspatful, dgene, embase, wpids COST IN U.S. DOLLARS

SINCE FILE TOTAL SESSION ENTRY 0.21 0.21

FULL ESTIMATED COST

FILE 'MEDLINE' ENTERED AT 13:52:47 ON 06 FEB 2004

FILE 'USPATFULL' ENTERED AT 13:52:47 ON 06 FEB 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE 'WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004 COPYRIGHT (C) 2004 THOMSON DERWENT

=> s albumin fusion proteins

2835 ALBUMIN FUSION PROTEINS

=> s cerebus protein

1 CEREBUS PROTEIN

=> s l1 and l2

0 L1 AND L2

=> s (cerebus protein) and albumin

O (CEREBUS PROTEIN) AND ALBUMIN

=> s 12 and fusion

0 L2 AND FUSION

=> d l2 ti abs ibib tot

ANSWER 1 OF 1 WPIDS COPYRIGHT 2004 THOMSON DERWENT on STN L2

Human and murine cerebus-like proteins - used for treating tissue defects TΙ and degenerative nerve conditions.

ΔN 1999-106054 [09] WPIDS

CR 2003-298696 [29]

9901553 A UPAB: 20030505 AR

A novel isolated DNA sequence comprises a DNA sequence selected from: (a) nucleotides beginning at # 1, 52, 55, 58, 61, 64, 67, 70, 73, 121, 256, 259, 262, 265, 268, 171, or 484 and ending at # 723 or 801 of the 804 bp DNA sequence given in the specification; and (b) sequences which hybridise to (a) under stringent hybridisation conditions and encode a protein which exhibits cerebus activity. Also claimed are: (1) an isolated DNA sequence comprising nucleotides encoding amino acids beginning at #1, 18 to 25, 41, 85 to 91 or 152, and ending at #241 or 267 of the 267 amino acid sequence given in the specification; (2) a vector comprising either of the above DNA molecules in operative association with an expression control sequence; (3) an isolated DNA molecule comprising nucleotides 268-801 of the 272 amino acid sequence given in the specification (sic), or naturally occurring allelic sequences of it; (4) a vector comprising the DNA of (4) in operative association with an expression control sequence; (5) an isolated DNA molecule encoding mammalian cerebus protein , comprising nucleotides 268-801 of the 804 bp DNA sequence given in the specification; (6) a vector comprising the DNA of (5) in operative

association with an expression control sequence; (7) a host cell

transformed with the vector of (2), (4) or (6); (8) a purified mammalian cerebus protein comprising the 267 amino acid sequence given in the specification; (9) a purified mammalian cerebus protein comprising residues 90-267 of the 272 amino acid sequence given in the specification; and (10) antibodies to the cerebus protein of (8) or (9).

USE - The host cell of (7) can be used to produce the mammalian cerebus proteins (claimed). Compositions containing the protein can be used in the formation of neurons and related neural cells and tissues, such as Schwann cells, glial cells, and astrocytes, as well as liver, pancreas, lung, heart, kidney, spleen, stomach, and cardiac tissue and cells. They may also be used to treat precursor or stem cells. The compositions can also be used for treating tissue defects, and healing and maintenance of various types of tissues and wounds. The mammalian cerebus protein containing compositions may also be used to treat or prevent degenerate nerve conditions such as Parkinson's disease, Alzheimer's disease, and Lou Gehrig's disease. They can also be used to treat osteoporosis, rheumatoid arthritis, osteoarthritis, and other abnormalities of connective tissue, or of other organs or tissues.

ACCESSION NUMBER:

1999-106054 [09] WPIDS

CROSS REFERENCE:

2003-298696 [29]

DOC. NO. CPI:

C1999-031758

TITLE:

Human and murine cerebus-like proteins - used for treating tissue defects and degenerative nerve

conditions.

B04 D16

DERWENT CLASS: INVENTOR(S):

DEROBERTIS, E M; FOLLETTIE, M

PATENT ASSIGNEE(S):

(GEMY) GENETICS INST INC; (REGC) UNIV CALIFORNIA

COUNTRY COUNT:

83

PATENT INFORMATION:

| PATENT NO | KIND DATE | WEEK | LA PG |
|-----------|-----------|------|-------|
|           | <b></b> - |      |       |

WO 9901553 A1 19990114 (199909) \* EN 50

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW

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57

AU 9878140 A 19990125 (199923)

US 5935852 A 19990810 (199938)

EP 1012278 A1 20000628 (200035) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

MX 2000000242 A1 20010601 (200235)

JP 2002511762 W 20020416 (200242)

AU 749031 B 20020620 (200252)

#### APPLICATION DETAILS:

| PAT      | TENT NO K                                | IND                | API            | PLICATION  | DATE   |
|----------|--|--------------------|----------------|--|--|
| AU<br>US | 9901553<br>9878140<br>5935852<br>1012278 | A1<br>A<br>A<br>A1 | AU<br>US<br>EP | 1998-US11462<br>1998-78140<br>1997-887997<br>1998-926263<br>1998-US11462 | 19980603<br>19980603<br>19970703<br>19980603<br>19980603 |
| JР       | 2000000242<br>2002511762<br>749031       |                    | MX<br>WO<br>JP | 1998-US11462<br>2000-242<br>1998-US11462<br>1999-507147<br>1998-78140    | 20000105<br>19980603<br>19980603<br>19980603             |

FILING DETAILS:

| PATENT NO K  | IND  | PATENT NO  |
|--|--|--|
| AU 9878140<br>EP 1012278<br>JP 2002511762<br>AU 749031 | A Based on<br>A1 Based on<br>W Based on<br>B Previous Publ<br>Based on | WO 9901553<br>WO 9901553<br>WO 9901553<br>. AU 9878140<br>WO 9901553 |

PRIORITY APPLN. INFO: US 1997-887997 19970703

=> file biosis

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 20.32 20.53

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004 COPYRIGHT (C) 2004 BIOLOGICAL ABSTRACTS INC.(R)

FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 4 February 2004 (20040204/ED)

FILE RELOADED: 19 October 2003.

=> d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN

L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

=> s 12

0 CEREBUS

1361492 PROTEIN

L6 0 CEREBUS PROTEIN

(CEREBUS (W) PROTEIN)

=> file medline, uspatful, dgene, embase, wpids, biosis, japio, fsta, jicst COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST 0.85 21.38

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FILE 'FSTA' ENTERED AT 14:00:26 ON 06 FEB 2004 COPYRIGHT (C) 2004 International Food Information Service

FILE 'JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004 COPYRIGHT (C) 2004 Japan Science and Technology Agency (JST)

=> s 12

L7 1 L2

=> d 17 ti abs ibib tot

L7 ANSWER 1 OF 1 WPIDS COPYRIGHT 2004 THOMSON DERWENT ON STN

TI Human and murine cerebus-like proteins - used for treating tissue defects and degenerative nerve conditions.

AN 1999-106054 [09] WPIDS

CR 2003-298696 [29]

AB WO 9901553 A UPAB: 20030505

A novel isolated DNA sequence comprises a DNA sequence selected from: (a) nucleotides beginning at # 1, 52, 55, 58, 61, 64, 67, 70, 73, 121, 256, 259, 262, 265, 268, 171, or 484 and ending at # 723 or 801 of the 804 bp DNA sequence given in the specification; and (b) sequences which hybridise to (a) under stringent hybridisation conditions and encode a protein which exhibits cerebus activity. Also claimed are: (1) an isolated DNA sequence comprising nucleotides encoding amino acids beginning at #1, 18 to 25, 41, 85 to 91 or 152, and ending at #241 or 267 of the 267 amino acid sequence given in the specification; (2) a vector comprising either of the above DNA molecules in operative association with an expression control sequence; (3) an isolated DNA molecule comprising nucleotides 268-801 of the 272 amino acid sequence given in the specification (sic), or naturally occurring allelic sequences of it; (4) a vector comprising the DNA of (4) in operative association with an expression control sequence; (5) an isolated DNA molecule encoding mammalian cerebus protein , comprising nucleotides 268-801 of the 804 bp DNA sequence given in the specification; (6) a vector comprising the DNA of (5) in operative association with an expression control sequence; (7) a host cell transformed with the vector of (2), (4) or (6); (8) a purified mammalian cerebus protein comprising the 267 amino acid sequence given in the specification; (9) a purified mammalian cerebus protein comprising residues 90-267 of the 272 amino acid sequence given in the specification; and (10) antibodies to the cerebus protein of (8) or (9).

USE - The host cell of (7) can be used to produce the mammalian cerebus proteins (claimed). Compositions containing the protein can be used in the formation of neurons and related neural cells and tissues, such as Schwann cells, glial cells, and astrocytes, as well as liver, pancreas, lung, heart, kidney, spleen, stomach, and cardiac tissue and cells. They may also be used to treat precursor or stem cells. The compositions can also be used for treating tissue defects, and healing and maintenance of various types of tissues and wounds. The mammalian cerebus protein containing compositions may also be used to treat or prevent degenerate nerve conditions such as Parkinson's disease, Alzheimer's disease, and Lou Gehrig's disease. They can also be used to treat osteoporosis, rheumatoid arthritis, osteoarthritis, and other abnormalities of connective tissue, or of other organs or tissues. Dwg.0/0

ACCESSION NUMBER:

WPIDS 1999-106054 [09]

CROSS REFERENCE:

2003-298696 [29]

DOC. NO. CPI:

C1999-031758

TITLE:

Human and murine cerebus-like proteins - used for

treating tissue defects and degenerative nerve

conditions.

DERWENT CLASS:

B04 D16

INVENTOR(S):

DEROBERTIS, E M; FOLLETTIE, M

PATENT ASSIGNEE(S):

(GEMY) GENETICS INST INC; (REGC) UNIV CALIFORNIA

COUNTRY COUNT: 83

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG\_\_\_\_\_\_

WO 9901553 A1 19990114 (199909) \* EN 50

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL

OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GE

GH GM GW HU ID IL IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG

57

MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG

UZ VN YU ZW

AU 9878140 A 19990125 (199923)

A 19990810 (199938) US 5935852

A1 20000628 (200035) EN EP 1012278

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

MX 2000000242 A1 20010601 (200235)

JP 2002511762 W 20020416 (200242)

AU 749031 B 20020620 (200252)

#### APPLICATION DETAILS:

| PATENT NO K   | IND | API | PLICATION    | DATE     |
|---------------|-----|-----|--------------|----------|
| WO 9901553    | A1  | WO. | 1998-US11462 | 19980603 |
| AU 9878140    | A   | ΑU  | 1998-78140   | 19980603 |
| US 5935852    | A   | US  | 1997-887997  | 19970703 |
| EP 1012278    | A1  | ΕP  | 1998-926263  | 19980603 |
|               |     | WO  | 1998-US11462 | 19980603 |
| MX 2000000242 | A1  | MX  | 2000-242     | 20000105 |
| JP 2002511762 | W   | WO  | 1998-US11462 | 19980603 |
|               |     | JP  | 1999-507147  | 19980603 |
| AU 749031     | В   | ΑU  | 1998-78140   | 19980603 |

#### FILING DETAILS:

| PATENT NO K  | IND     |          |       | PA             | rent no                                  | _ |
|--|---------|----------|-------|----------------|--|---|
| AU 9878140<br>EP 1012278<br>JP 2002511762<br>AU 749031 | A1<br>W | Previous | Publ. | WO<br>WO<br>AU | 9901553<br>9901553<br>9901553<br>9878140 | _ |
|  |         | Based on |       | WU             | 9901553                                  |   |

PRIORITY APPLN. INFO: US 1997-887997 19970703

## => d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L21 S CEREBUS PROTEIN L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

L6 0 S L2

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

L7 1 S L2

=> s TIMP-1 or tissue inhibitor metalloproteinase-1

5 FILES SEARCHED...

L8 8080 TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1

=> s 18 and 11

L9 5 L8 AND L1

=> d 19 ti abs ibib tot

L9 ANSWER 1 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL Albumin fusion proteins

INVENTOR(S):

TITLE:

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

| NUMBER                            |          | DATE                 |     |
|-----------------------------------|----------|----------------------|-----|
| <br>S 2004010134<br>S 2001-833245 | A1<br>A1 | 20040115<br>20010412 | (9) |

Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18

18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 2 OF 5 USPATFULL on STN

TI Albumin fusion proteins

AB The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic

acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|  | NUMBER | KIND | DATE |     |
|--|--------|------|------|-----|
| PATENT INFORMATION: US 2003219875 A1 20031127 APPLICATION INFO.: US 2001-833118 A1 20010412 (9 | <br>   | ***  |      | (9) |

NUMBER DATE

PRIORITY INFORMATION:

US 2000-256931P 20001221 (60) US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: FILE SEGMENT: Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 29

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L9 ANSWER 3 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL Albumin fusion proteins

INVENTOR (S):

TITLE:

Ballance, David J., Berwyn, PA, UNITED STATES Sleep, Darrell, West Bridgford, UNITED KINGDOM Prior, Christopher P., Rosemont, PA, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

| •                   | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003199043  | A1   | 20031023 |     |
| APPLICATION INFO.:  | ÚS 2001-832501 | A1   | 20010412 | (9) |

NUMBER DATE

PRIORITY INFORMATION:

US 2000-256931P

20001221 (60)

20000425 (60) US 2000-199384P 20000412 (60) US 2000-229358P

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 4 OF 5 USPATFULL on STN 1.9

Albumin fusion proteins TΙ

The present invention encompasses albumin fusion proteins. Nucleic acid AB molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:244853 USPATFULL Albumin fusion proteins

TITLE: INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND | DATE      |     |
|---------------------|----------------|------|-----------|-----|
|                     |                |      | - <b></b> |     |
| PATENT INFORMATION: | US 2003171267  | A1   | 20030911  |     |
| APPLICATION INFO.:  | US 2001-833117 | A1   | 20010412  | (9) |

|          |              |    | NUMBER                                       | DATE                             |      |
|----------|--------------|----|--|----------------------------------|------|
| PRIORITY | INFORMATION: | US | 2000-256931P<br>2000-199384P<br>2000-229358P | 20001221<br>20000425<br>20000412 | (60) |

DOCUMENT TYPE: FILE SEGMENT:

Utility

LEGAL REPRESENTATIVE:

APPLICATION

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

59

NUMBER OF DRAWINGS:

20 Drawing Page(s)

LINE COUNT:

13208

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- ANSWER 5 OF 5 USPATFULL on STN L9
- Albumin fusion proteins ΤI
- The present invention encompasses albumin fusion proteins. Nucleic acid AB molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion

proteins of the invention.

L12 ANSWER 1 OF 5 USPATFULL on STN Albumin fusion proteins

The present invention encompasses albumin fusion

ΤI

AB

CAS INDEXING IS AVAILABLE FOR THIS PATENT. 2003:181414 USPATFULL ACCESSION NUMBER: Albumin fusion proteins TITLE: Rosen, Craig A., Laytonsville, MD, UNITED STATES INVENTOR(S): Haseltine, William A., Washington, DC, UNITED STATES KIND DATE NUMBER US 2003125247 A1 20030703 US 2001-833041 A1 20010412 PATENT INFORMATION: APPLICATION INFO.: DATE NUMBER \_\_\_\_\_ US 2000-256931P 20001221 (60) PRIORITY INFORMATION: US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60) Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE: ROCKVILLE, MD, 20850 NUMBER OF CLAIMS: 29 EXEMPLARY CLAIM: 1 NUMBER OF DRAWINGS: 20 Drawing Page(s) 15235 LINE COUNT: CAS INDEXING IS AVAILABLE FOR THIS PATENT. => d his (FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004) FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004 2835 S ALBUMIN FUSION PROTEINS L11 S CEREBUS PROTEIN L2L3 0 S L1 AND L2 0 S (CEREBUS PROTEIN) AND ALBUMIN L40 S L2 AND FUSION L5 FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004 L6 0 S L2 FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004 L7 1 S L2 8080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1 L8L9 5 S L8 AND L1 => s 18 and fusion 378 L8 AND FUSION L10 => s 110 and albumin 221 L10 AND ALBUMIN => s l11 and albumin fragment 5 L11 AND ALBUMIN FRAGMENT => d l12 ti abs ibib tot

proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Haseltine, William A., Washington, DC, UNITED STATES

|  | NUMBER                          | KIND     | DATE                 |     |
|--|---------------------------------|----------|----------------------|-----|
| PATENT INFORMATION: APPLICATION INFO.: | US 2004010134<br>US 2001-833245 | A1<br>A1 | 20040115<br>20010412 | (9) |

NUMBER DATE \_\_\_\_\_\_

PRIORITY INFORMATION:

US 2000-256931P 20001221 (60)

US 2000-199384P 20000425 (60)

US 2000-229358P 20000412 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

# L12 ANSWER 2 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion AB proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                       | NUMBER        | KIND | DATE     |     |
|-----------------------|---------------|------|----------|-----|
| -                     |               |      |          |     |
| PATENT INFORMATION: U | S 2003219875  | A1   | 20031127 |     |
| APPLICATION INFO.: U  | S 2001-833118 | A1   | 20010412 | (9) |

NUMBER DATE PRIORITY INFORMATION: US 2000-256931P 20001221 (60)

US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility
FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 25 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### L12 ANSWER 3 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL Albumin fusion proteins

TITLE: INVENTOR(S):

Ballance, David J., Berwyn, PA, UNITED STATES
Sleep, Darrell, West Bridgford, UNITED KINGDOM
Prior, Christopher P., Rosemont, PA, UNITED STATES
Sadeghi, Homayoun, Doylestown, PA, UNITED STATES
Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND      | DATE     |     |
|---------------------|----------------|-----------|----------|-----|
|                     |                | <b></b> - |          |     |
| PATENT INFORMATION: | US 2003199043  | A1        | 20031023 |     |
| APPLICATION INFO .: | US 2001-832501 | A1        | 20010412 | (9) |

NUMBER DATE
-----PRIORITY INFORMATION: US 2000-256931P 20001221 (60)
US 2000-199384P 20000425 (60)
US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 60 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 14339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## L12 ANSWER 4 OF 5 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using

these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:244853 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003171267  | A1   | 20030911 |     |
| APPLICATION INFO.:  | US 2001-833117 | A1   | 20010412 | (9) |

NUMBER DATE \_\_\_\_\_ \_\_\_ ÚS 2000-256931P 20001221 (60) PRIORITY INFORMATION: US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

59

NUMBER OF DRAWINGS:

20 Drawing Page(s)

LINE COUNT:

13208

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L12 ANSWER 5 OF 5 USPATFULL on STN

Albumin fusion proteins TIAB

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:181414 USPATFULL

TTTLE

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

20000412 (60)

|  | NUMBER                             | KIND                           | DATE                 |     |
|--|------------------------------------|--------------------------------|----------------------|-----|
| PATENT INFORMATION: APPLICATION INFO : | US 2003125247<br>US 2001-833041    | A1<br>A1                       | 20030703<br>20010412 | (9) |
|  | NUMBER                             | DATE                           |                      |     |
| PRIORITY INFORMATION:                  | US 2000-256931P<br>US 2000-199384P | 20001221 (60)<br>20000425 (60) |                      |     |

US 2000-229358P

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 29

NUMBER OF DRAWINGS:

20 Drawing Page(s)

LINE COUNT:

15235

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### => d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN

L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

L6 0 S L2

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

L7 1 S L2

L8 8080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1

L9 5 S L8 AND L1

L10 378 S L8 AND FUSION

L11 . 221 S L10 AND ALBUMIN

L12 5 S L11 AND ALBUMIN FRAGMENT

=> s l11 and shelf-life

L13 9 L11 AND SHELF-LIFE

## => d l13 ti abs ibib tot

L13 ANSWER 1 OF 9 USPATFULL on STN

TI Biospecific contrast agents

Methods and apparatuses for detecting a condition of a sample (including cervical cancers and pre-cancers) through reflectance and/or fluorescence imaging. A sample is obtained. One or more metallic nanoparticles and/or one or more quantum dots are obtained. The one or more metallic nanoparticles and/or one or more quantum dots are coupled to one or more biomarkers of the sample that are associated with the condition. A reflectance and/or fluorescence image of the sample is then taken. The image(s) exhibit characteristic optical scattering from the one or more metallic nanoparticles and/or characteristic fluorescence excitation from the one or more quantum dots to signal the presence of the one or more biomarkers. In this way, the condition can be readily screened or diagnosed.

ACCESSION NUMBER:

2004:31276 USPATFULL

TITLE:

Biospecific contrast agents

INVENTOR(S):

Sokolov, Konstantin, Austin, TX, UNITED STATES
Korgel, Brian A., Round Rock, TX, UNITED STATES
Ellington, Andrew D., Austin, TX, UNITED STATES
Richards-Kortum, Rebecca, Austin, TX, UNITED STATES

NUMBER KIND DATE

\_\_\_\_\_

PATENT INFORMATION:

A1 US 2004023415

20040205 A1

APPLICATION INFO .:

US 2003-382136

20030305 (10)

DATE NUMBER \_\_\_\_\_\_\_

PRIORITY INFORMATION:

US 2002-361924P

20020305 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

Michael C. Barrett, Esq., FULBRIGHT & JAWORSKI, L.L.P.,

600 Congress Avenue, Suite 2400, Austin, TX, 78701

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

11 Drawing Page(s)

LINE COUNT:

3948

## L13 ANSWER 2 OF 9 USPATFULL on STN

Albumin fusion proteins TI

AB

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2004010134  | A1   | 20040115 |     |
| APPLICATION INFO.:  | US 2001-833245 | A1   | 20010412 | (9) |

|          |              |    | NUMBER                                       | DATE                             |      |
|----------|--------------|----|--|----------------------------------|------|
| PRIORITY | INFORMATION: | US | 2000-256931P<br>2000-199384P<br>2000-229358P | 20001221<br>20000425<br>20000412 | (60) |

Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

29 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

## L13 ANSWER 3 OF 9 USPATFULL on STN

Nanoporous particle with a retained target TI

Porous nanostructured materials, such as porous nanostructured liquid AΒ and liquid crystalline particles or materials, incorporate a target substantially within the material which selectively binds a chemical of interest which can diffusion within the porous nanostructured material and be bound by the target. The porous nanostructured materials can be dispersed as particles in a medium in which said chemical of interest is located with low turbidity. Markers which detect binding of said

chemical of interest can be maintained in the medium separate and apart from the target, and any active compound (e.g., an enzyme) associated therewith by the porous nanostructured material, such that detectable changes in the marker only result when the active compounds diffuse out of the porous nanostructured materials after the chemical of interest binds to the target.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:330129 USPATFULL

TITLE:

Nanoporous particle with a retained target

INVENTOR(S):

Anderson, David, Colonial Heights, VA, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_\_ US 2003232340 A1 20031218

PATENT INFORMATION:

APPLICATION INFO.:

US 2002-170214

A1 20020613 (10)

DOCUMENT TYPE: FILE SEGMENT:

Utility

LEGAL REPRESENTATIVE:

APPLICATION WHITHAM, CURTIS & CHRISTOFFERSON, P.C., 11491 SUNSET

HILLS ROAD, SUITE 340, RESTON, VA, 20190

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

2 Drawing Page(s)

LINE COUNT:

2555

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 4 OF 9 USPATFULL on STN

Albumin fusion proteins ΤI

The present invention encompasses albumin fusion AΒ

proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating,

preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:312278 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|                     | NUMBER         | KIND | DATE     |     |
|---------------------|----------------|------|----------|-----|
|                     |                |      |          |     |
| PATENT INFORMATION: | US 2003219875  | A1   | 20031127 |     |
| APPLICATION INFO.:  | US 2001-833118 | A1   | 20010412 | (9) |

NUMBER DATE

PRIORITY INFORMATION:

US 2000-256931P 20001221 (60)

US 2000-199384P 20000425 (60) US 2000-229358P 20000412 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

29

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

15415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 5 OF 9 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:282700 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR (S):

Ballance, David J., Berwyn, PA, UNITED STATES Sleep, Darrell, West Bridgford, UNITED KINGDOM Prior, Christopher P., Rosemont, PA, UNITED STATES Sadeghi, Homayoun, Doylestown, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|   | NUMBER                          | KIND     | DATE                 |     |
|---|---------------------------------|----------|----------------------|-----|
| PATENT INFORMATION:<br>APPLICATION INFO.: | US 2003199043<br>US 2001-832501 | A1<br>A1 | 20031023<br>20010412 | (9) |
|   | NUMBER                          | DA       | TE                   |     |
|   |                                 |          |                      |     |

PRIORITY INFORMATION: US 2000-256931P 20001221 (60)
US 2000-199384P 20000425 (60)
US 2000-229358P 20000412 (60)

DOCUMENT TYPE: FILE SEGMENT: Utility APPLICATION

FILE SEGMENT: APPLICATI

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS:

1

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

18 Drawing Page(s)

LINE COUNT:

14339

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L13 ANSWER 6 OF 9 USPATFULL on STN

TI Albumin fusion proteins

The present invention encompasses albumin fusion proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2

2003:244853 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Sadeghi, Homayoun, Doylestown, PA, UNITED STATES

Prior, Christopher P., Rosemont, PA, UNITED STATES Turner, Andrew J., Eagleville, PA, UNITED STATES

|  | NUMBER  | KIND  | DATE       |                  |
|--|---|-------|------------|------------------|
|  | US 2003171267<br>US 2001-833117                         |       |            | (9)              |
|  | NUMBER  | DAT   | ΓE         |                  |
| PRIORITY INFORMATION:  | US 2000-256931P<br>US 2000-199384P<br>US 2000-229358P   | 20000 | 0425 (60)  |                  |
| DOCUMENT TYPE: FILE SEGMENT: LEGAL REPRESENTATIVE:   | Utility APPLICATION HUMAN GENOME SCIE ROCKVILLE, MD, 20 |       | NC, 9410 F | KEY WEST AVENUE, |
| NUMBER OF CLAIMS: EXEMPLARY CLAIM: NUMBER OF DRAWINGS: LINE COUNT: CAS INDEXING IS AVAILAB | 13208   |       |            |                  |

L13 ANSWER 7 OF 9 USPATFULL on STN

Albumin fusion proteins TI

The present invention encompasses albumin fusion AB proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:181414 USPATFULL

TITLE:

Albumin fusion proteins

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

|  | NUMBER  | KIND | DATE      |                  |
|--|---|------|-----------|------------------|
| PATENT INFORMATION: APPLICATION INFO.:           |   |      |           |                  |
|  | NUMBER  | DA   | TE        |                  |
| PRIORITY INFORMATION:                            | US 2000-256931P<br>US 2000-199384P<br>US 2000-229358P | 2000 | 0425 (60) |                  |
| DOCUMENT TYPE: FILE SEGMENT:                     | Utility<br>APPLICATION                                |      |           |                  |
| LEGAL REPRESENTATIVE:                            |   |      | NC, 9410  | KEY WEST AVENUE, |
| NUMBER OF CLAIMS:                                | 29<br>1   |      |           |                  |
| EXEMPLARY CLAIM: NUMBER OF DRAWINGS: LINE COUNT: |   | )    |           |                  |
| CAS INDEXING IS AVAILAB                          | LE FOR THIS PATENT                                    | ·•   |           |                  |

L13 ANSWER 8 OF 9 USPATFULL on STN

Coated particles, methods of making and using ΤI

A particle coated with a nonlamellar material such as a nonlamellar AΒ crystalline material, a nonlamellar amorphous material, or a nonlamellar semi-crystalline material includes an internal matrix core having at least one a nanostructured liquid phase, or at least on nanostructured liquid crystalline phase or a combination of the two is used for the delivery of active agents such as pharmaceuticals, nutrients, pesticides, etc. The coated particle can be fabricated by a variety of different techniques where the exterior coating is a nonlamellar material such as a nonlamellar crystalline material, a nonlamellar amorphous material, or a nonlamellar semi-crystalline material

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:159130 USPATFULL

TITLE:

Coated particles, methods of making and using

INVENTOR(S):

Anderson, David M., Colonial Heights, VA, UNITED STATES

NUMBER KIND DATE \_\_\_\_\_\_ US 2003108743 A1 US 6638621 B2 US 2002-170237 A1 PATENT INFORMATION: 20030612 20031028 (10)

APPLICATION INFO.: 20020613

Continuation-in-part of Ser. No. US 2000-297997, filed RELATED APPLN. INFO.:

on 16 Aug 2000, GRANTED, Pat. No. US 6482517

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

WHITHAM, CURTIS & CHRISTOFFERSON, P.C., 11491 SUNSET LEGAL REPRESENTATIVE:

HILLS ROAD, SUITE 340, RESTON, VA, 20190

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 11 Drawing Page(s)

LINE COUNT: 5538

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 9 OF 9 USPATFULL on STN

TΙ Multifunctional protease inhibitors and their use in treatment of

Fusion proteins of protease inhibitors are provided, in AB particular fusion proteins of alpha 1-antitrypsin (AAT) and a second protease inhibitor, such as secretory leukocyte protease inhibitor (SLPI) or tissue inhibitor of metalloproteases (TIMP). Polynucleotides encoding the fusion proteins, vectors comprising such polynucleotides, and host cells containing such vectors are also provided. Methods of making the fusion proteins of the invention are also provide, as well as methods of using the fusion proteins, for example to inhibit protease activity in a biological sample or in the treatment of an individual suffering from, or at risk for, a disease or disorder involving unwanted protease activity.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:106306 USPATFULL

TITLE:

Multifunctional protease inhibitors and their use in

treatment of disease

INVENTOR(S):

Barr, Philip J., Oakland, CA, UNITED STATES Gibson, Helen, Oakland, CA, UNITED STATES

Pemberton, Philip, San Francisco, CA, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_\_ US 2003073217 A1 20030417 PATENT INFORMATION: APPLICATION INFO.: US 2001-25514 A1 20011218 (10)

> NUMBER DATE

\_\_\_\_\_

PRIORITY INFORMATION: US 2000-256699P 20001218 (60) US 2001-331966P 20011120 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: MORRISON & FOERSTER LLP, 755 PAGE MILL RD, PALO ALTO,

CA, 94304-1018

NUMBER OF CLAIMS: 35 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 6 Drawing Page(s)

LINE COUNT: 3252

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

### => d his

(FILE 'HOME' ENTERED AT 13:52:01 ON 06 FEB 2004)

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS' ENTERED AT 13:52:47 ON 06 FEB 2004

L1 2835 S ALBUMIN FUSION PROTEINS

L2 1 S CEREBUS PROTEIN

L3 0 S L1 AND L2

L4 0 S (CEREBUS PROTEIN) AND ALBUMIN

L5 0 S L2 AND FUSION

FILE 'BIOSIS' ENTERED AT 13:59:37 ON 06 FEB 2004

L6 0 S L2

FILE 'MEDLINE, USPATFULL, DGENE, EMBASE, WPIDS, BIOSIS, JAPIO, FSTA, JICST-EPLUS' ENTERED AT 14:00:26 ON 06 FEB 2004

L7 1 S L2

L8 8080 S TIMP-1 OR TISSUE INHIBITOR METALLOPROTEINASE-1

L9 5 S L8 AND L1

L10 378 S L8 AND FUSION L11 221 S L10 AND ALBUMIN

L12 5 S L11 AND ALBUMIN FRAGMENT

L13 9 S L11 AND SHELF-LIFE

=> s lll and N-terminus fusion

L14 0 L11 AND N-TERMINUS FUSION

=> s l11 and C-terminus fusion

L15 0 L11 AND C-TERMINUS FUSION

=> d l11 ti abs ibib 1-25

L11 ANSWER 1 OF 221 USPATFULL on STN

TI Biospecific contrast agents

Methods and apparatuses for detecting a condition of a sample (including cervical cancers and pre-cancers) through reflectance and/or fluorescence imaging. A sample is obtained. One or more metallic nanoparticles and/or one or more quantum dots are obtained. The one or more metallic nanoparticles and/or one or more quantum dots are coupled to one or more biomarkers of the sample that are associated with the condition. A reflectance and/or fluorescence image of the sample is then taken. The image(s) exhibit characteristic optical scattering from the one or more metallic nanoparticles and/or characteristic fluorescence excitation from the one or more quantum dots to signal the presence of the one or more biomarkers. In this way, the condition can be readily screened or diagnosed.

ACCESSION NUMBER:

2004:31276 USPATFULL

TITLE:

Biospecific contrast agents

INVENTOR(S):

Sokolov, Konstantin, Austin, TX, UNITED STATES Korgel, Brian A., Round Rock, TX, UNITED STATES Ellington, Andrew D., Austin, TX, UNITED STATES Richards-Kortum, Rebecca, Austin, TX, UNITED STATES

KIND DATE NUMBER \_\_\_\_\_\_ US 2004023415 A1 US 2003-382136 A1 20040205 20030305 (10)

NUMBER DATE

\_\_\_\_\_

PRIORITY INFORMATION:

PATENT INFORMATION:

APPLICATION INFO .:

US 2002-361924P 20020305 (60) Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE: Michael C. Barrett, Esq., FULBRIGHT & JAWORSKI, L.L.P.,

600 Congress Avenue, Suite 2400, Austin, TX, 78701

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 11 Drawing Page(s)

LINE COUNT: 3948

L11 ANSWER 2 OF 221 USPATFULL on STN

Biochips for characterizing biological processes ΤI

This invention includes biochips for analysis of a variety of molecules, AB cell components and cells. Embodiments of this invention include devices and methods for the parallel and/or nearly parallel processing of biological analytes. Biochips can comprise a substrate, Raman signal-enhancing structures, and receptors selective and/or specific for the analyte(s) to be assayed. Biochips can be read using a Raman reader and can provide for rapid, sensitive, direct assays for physiological and/or pathophysiological conditions of interest.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:31155 USPATFULL

TITLE: INVENTOR(S): Biochips for characterizing biological processes Kreimer, David I., Berkeley, CA, UNITED STATES Nufert, Thomas H., Walnut Creek, CA, UNITED STATES

Ginzburg, Lev, Fremont, CA, UNITED STATES Yevin, Oleg A., Oakland, CA, UNITED STATES

DATE NUMBER KIND

-----US 2004023293 A1 20040205 US 2002-294385 A1 20021114 (10) PATENT INFORMATION: APPLICATION INFO.:

Continuation-in-part of Ser. No. US 2001-925189, filed RELATED APPLN. INFO.:

on 8 Aug 2001, PENDING Continuation-in-part of Ser. No.

US 2001-815909, filed on 23 Mar 2001, PENDING

Continuation-in-part of Ser. No. US 2000-670453, filed

on 26 Sep 2000, PENDING

DATE NUMBER \_\_\_\_\_

US 1999-156195P 19990927 (60) PRIORITY INFORMATION: US 2001-336445P 20011114 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: Sheldon R. Meyer, FLIESLER DUBB MEYER & LOVEJOY LLP, Fourth Floor, Four Embarcadero Center, San Francisco,

CA, 94111-4156

NUMBER OF CLAIMS: 40 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 37 Drawing Page(s)

LINE COUNT: 3572 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 3 OF 221 USPATFULL on STN

ΤI Proteases

The invention provides human proteases (PRTS) and polynucleotides which AB identify and encode PRTS. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for diagnosing, treating, or preventing disorders associated with aberrant expression of PRTS.

ACCESSION NUMBER:

2004:31105 USPATFULL

TITLE:

INVENTOR(S):

Proteases Henry, Yue, Sunnyvale, CA, UNITED STATES

Elliott, Vicki S, San Jose, CA, UNITED STATES R Gandhi, Ameena, San Francisco, CA, UNITED STATES Lal, Preeti G, Santa Clara, CA, UNITED STATES Au-Young, Janice, Brisbane, CA, UNITED STATES Tribouley, Catherine M, San Francisco, CA, UNITED

STATES Delegeane, Angelo M, Milpitas, CA, UNITED STATES Baughn, Mariah R, San Leandro, CA, UNITED STATES Nguyen, Danniel B, San Jose, CA, UNITED STATES Lee, Ernestine A, Albany, CA, UNITED STATES Hafalia, April J A, Daly City, CA, UNITED STATES Khan, Farrah A, Des Plaines, IL, UNITED STATES Chawla, Narinder K, Union City, CA, UNITED STATES Yao, Monique G, Carmel, IN, UNITED STATES Lu, Dyung Aina M, San Jose, CA, UNITED STATES Arvizu, Chandra S, San Jose, CA, UNITED STATES Tang, Y Tom, San Jose, CA, UNITED STATES Walsh, Roderick T, Canterbury, UNITED KINGDOM Azimzai, Yalda, Oakland, CA, UNITED STATES Lu, Yan, Palo Alto, CA, UNITED STATES Ramkumar, Jayalaxmi, Fremont, CA, UNITED STATES Xu, Yuming, Mountain View, CA, UNITED STATES Reddy, Roopa, Sunnyvale, CA, UNITED STATES Das, Debopriya, Mountain View, CA, UNITED STATES Kearney, Liam, San Francisco, CA, UNITED STATES

Kallick, Deborah A, Galveston, TX, UNITED STATES

|    | NUMBER      | KIND | DATE     |
|----|-------------|------|----------|
|    |             |      |          |
| US | 2004023243  | A1   | 20040205 |
| US | 2003-311035 | A1   | 20030519 |

PATENT INFORMATION: APPLICATION INFO.:

DOCUMENT TYPE:

WO 2001-US19178 Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

INCYTE CORPORATION (formerly known as Incyte, Genomics,

20010613

(10)

Inc.), 3160 PORTER DRIVE, PALO ALTO, CA, 94304

116 NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 8891 LINE COUNT:

ANSWER 4 OF 221 USPATFULL on STN L11

Novel human gene relating to respiratory diseases, obesity, and TI inflammatory bowel disease

This invention relates to genes identified from human chromosome AΒ 20p13-p12, which are associated with various diseases, including asthma. The invention also relates to the nucleotide sequences of these genes, isolated nucleic acids comprising these nucleotide sequences, and isolated polypeptides or peptides encoded thereby. The invention further relates to vectors and host cells comprising the disclosed nucleotide sequences, or fragments thereof, as well as antibodies that bind to the encoded polypeptides or peptides. Also related are ligands that modulate the activity of the disclosed genes or gene products. In addition, the invention relates to methods and compositions employing the disclosed nucleic acids, polypeptides or peptides, antibodies, and/or ligands for use in diagnostics and therapeutics for asthma and other diseases.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

INVENTOR(S):

2004:31077 USPATFULL

TITLE:

Novel human gene relating to respiratory diseases,

obesity, and inflammatory bowel disease

Keith, Tim, Bedford, MA, UNITED STATES

Little, Randall D., Newtonville, MA, UNITED STATES

Eerdewegh, Paul Van, Weston, MA, UNITED STATES

Dupuis, Josee, Newton, MA, UNITED STATES

Del Mastro, Richard G., Norfolk, MA, UNITED STATES

Simon, Jason, Westfield, NJ, UNITED STATES Allen, Kristina, Hopkinton, MA, UNITED STATES Pandit, Sunil, Gaithersburg, MD, UNITED STATES

NUMBER KIND DATE \_\_\_\_\_

PATENT INFORMATION: APPLICATION INFO.:

US 2004023215 A1 20040205 US 2002-126022 A1 20020419 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2001-834597, filed on 13 Apr 2001, PENDING Continuation-in-part of Ser. No. US 2000-548797, filed on 13 Apr 2000, PENDING

NUMBER DATE 

PRIORITY INFORMATION:

US 1999-129391P 19990413 (60)

DOCUMENT TYPE:

Utility. APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

MORGAN & FINNEGAN, L.L.P., 345 Park Avenue, New York,

NY, 10154-0053

NUMBER OF CLAIMS:

73

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

157 Drawing Page(s)

LINE COUNT:

20001

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 5 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ΤI

The present invention relates to novel proteins. More specifically, AB isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

ACCESSION NUMBER:

2004:25127 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR (S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES

NUMBER KIND DATE

PATENT INFORMATION:

US 2004018969

A1 20040129

| APPLICATION INFO.:    | US 2001-764875   | A1 20010117   | (9) |
|-----------------------|--|---|-----|
|                       | NUMBER   | DATE  |     |
| PRIORITY INFORMATION: | US 2000-179065P<br>US 2000-180628P<br>US 2000-214886P<br>US 2000-217487P<br>US 2000-225758P<br>US 2000-220963P                                       | 20000131 (60)<br>20000204 (60)<br>20000628 (60)<br>20000711 (60)<br>20000814 (60)<br>20000726 (60)                  |     |
|                       | US 2000-217496P<br>US 2000-225447P<br>US 2000-218290P<br>US 2000-225757P<br>US 2000-226868P<br>US 2000-216647P<br>US 2000-225267P                    | 20000711 (60)<br>20000814 (60)<br>20000714 (60)<br>20000814 (60)<br>20000822 (60)<br>20000707 (60)<br>20000814 (60) |     |
|                       | US 2000-216880P<br>US 2000-225270P<br>US 2000-251869P<br>US 2000-235834P<br>US 2000-234274P<br>US 2000-234223P<br>US 2000-228924P                    | 20000707 (60)<br>20000814 (60)<br>20001208 (60)<br>20000927 (60)<br>20000921 (60)<br>20000921 (60)<br>20000830 (60) |     |
|                       | US 2000-224518P<br>US 2000-236369P<br>US 2000-224519P<br>US 2000-220964P<br>US 2000-241809P<br>US 2000-249299P<br>US 2000-236327P                    | 20000814 (60)<br>20000929 (60)<br>20000814 (60)<br>20000726 (60)<br>20001020 (60)<br>20001117 (60)<br>20000929 (60) |     |
|                       | US 2000-241785P<br>US 2000-244617P<br>US 2000-225268P<br>US 2000-236368P<br>US 2000-251856P<br>US 2000-251868P                                       | 20001020 (60)<br>20001101 (60)<br>20000814 (60)<br>20000929 (60)<br>20001208 (60)<br>20001208 (60)                  |     |
|                       | US 2000-229344P<br>US 2000-234997P<br>US 2000-229343P<br>US 2000-229345P<br>US 2000-229287P<br>US 2000-229513P                                       | 20000901 (60)<br>20000925 (60)<br>20000901 (60)<br>20000901 (60)<br>20000905 (60)                                   |     |
|                       | US 2000-231413P<br>US 2000-229509P<br>US 2000-236367P<br>US 2000-237039P<br>US 2000-237038P<br>US 2000-236370P<br>US 2000-236802P                    | 2000908 (60)<br>20000905 (60)<br>20000929 (60)<br>20001002 (60)<br>20001002 (60)<br>20000929 (60)<br>20001002 (60)  |     |
|                       | US 2000-238802P<br>US 2000-237037P<br>US 2000-237040P<br>US 2000-240960P<br>US 2000-239935P<br>US 2000-239937P<br>US 2000-241787P                    | 20001002 (60)<br>20001002 (60)<br>20001002 (60)<br>20001020 (60)<br>20001013 (60)<br>20001013 (60)<br>20001020 (60) |     |
|                       | US 2000-246474P<br>US 2000-246532P<br>US 2000-249216P<br>US 2000-249210P<br>US 2000-226681P<br>US 2000-225759P<br>US 2000-225213P<br>US 2000-227182P | 20001108 (60)<br>20001108 (60)<br>20001117 (60)<br>20001117 (60)<br>20000822 (60)<br>20000814 (60)<br>20000822 (60) | ·   |

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US 2000-225214P
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                    20001211 (60)
US 2000-254097P
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US 2000-231968P 20000912 (60) 20000818 (60) US 2000-226279P 20000302 (60) US 2000-186350P 20000224 (60) US 2000-184664P 20000316 (60) US 2000-189874P 20000418 (60) US 2000-198123P US 2000-227009P 20000823 (60) 20000926 (60) US 2000-235484P 20000317 (60) US 2000-190076P 20000607 (60) US 2000-209467P US 2000-205515P 20000519 (60) US 2001-259678P 20010105 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM: LINE COUNT: 24 1 38235

L11 ANSWER 6 OF 221 USPATFULL on STN

TI Molecules for diagnostics and therapeutics

The present invention provides purified human polynucleotides for diagnostics and therapeutics (dithp). Also encompassed are the polypeptides (DITHP) encoded by dithp. The invention also provides for the use of dithp, or complements, oligonucleotides, or fragments thereof in diagnostic assays. The invention further provides for vectors and host cells containing dithp for the expression of DITHP. The invention additionally provides for the use of isolated and purified DITHP to induce antibodies and to screen libraries of compounds and the use of anti-DITHP antibodies in diagnostic assays. Also provided are microarrays containing dithp and methods of use.

ACCESSION NUMBER:

Τ,

INVENTOR(S):

TITLE:

2004:18785 USPATFULL

Molecules for diagnostics and therapeutics
Hodgson, David M., Ann Arbor, MI, UNITED STATES
Lincoln, Stephen E., Potomac, MD, UNITED STATES
Russo, Frank D., Sunnyvale, CA, UNITED STATES
Albany, Peter A., Berkeley, CA, UNITED STATES
Banville, Steve C., Sunnyvale, CA, UNITED STATES
Bratcher, Shawn R., Mountain View, CA, UNITED STATES
Dufour, Gerard E., Castro Valley, CA, UNITED STATES
Cohen, Howard J., Palo Alto, CA, UNITED STATES
Rosen, Bruce H., Menlo Park, CA, UNITED STATES
Chalup, Michael S., Livingston, TX, UNITED STATES
Jackson, Jennifer L., Santa Cruz, CA, UNITED STATES
Jones, Anissa L., San Jose, CA, UNITED STATES
Yu, Jimmy Y., Fremont, CA, UNITED STATES
Greenawalt, Lila B., San Jose, CA, UNITED STATES

Panzer, Scott R., Sunnyvale, CA, UNITED STATES Roseberry Lincoln, Ann M., Potomac, MD, UNITED STATES

Wright, Rachel J., Merivale, NEW ZEALAND

Daniels, Susan E., Mountain View, CA, UNITED STATES
Incyte Corporation, Palo Alto, CA, UNITED STATES (U.S.

corporation)

PATENT ASSIGNEE(S):

| NUMBER       | KIND | DATE     |
|--------------|------|----------|
|              |      |          |
| S 2004014087 | Α1   | 20040122 |

PATENT INFORMATION: APPLICATION INFO.: RELATED APPLN. INFO.:

US 2004014087 A1 20040122 US 2003-378029 A1 20030228 (10)

Continuation-in-part of Ser. No. US 2001-980285, filed on 30 Nov 2001, PENDING A 371 of International Ser. No. WO 2000-US15404, filed on 31 May 2000, PENDING

NUMBER DATE \_\_\_\_\_ 19990805 (60) US 1999-147500P PRIORITY INFORMATION: 19990805 (60) US 1999-147542P 19990805 (60) US 1999-147541P US 1999-147824P 19990805 (60) 19990805 (60) US 1999-147547P 19990805 (60) US 1999-147530P US 1999-147536P 19990805 (60) US 1999-147520P 19990805 (60) US 1999-147527P 19990805 (60) US 1999-147549P 19990805 (60) US 1999-147377P 19990804 (60) US 1999-147436P 19990804 (60) 19990603 (60) US 1999-137411P 19990603 (60) US 1999-137396P 19990603 (60) US 1999-137417P US 1999-137337P 19990603 (60) 19990602 (60) US 1999-137173P 19990602 (60) US 1999-137114P 19990602 (60) US 1999-137259P US 1999-137113P 19990602 (60) US 1999-137260P 19990602 (60) 19990602 (60) US 1999-137258P 19990602 (60) US 1999-137109P 19990601 (60) US 1999-137161P Utility

DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

INCYTE CORPORATION (formerly known as Incyte, Genomics,

Inc.), 3160 PORTER DRIVE, PALO ALTO, CA, 94304

NUMBER OF CLAIMS: 19
EXEMPLARY CLAIM: 1
LINE COUNT: 14819

L11 ANSWER 7 OF 221 USPATFULL on STN

TI Nucleic acids, proteins, and antibodies

The present invention relates to novel proteins. More specifically, isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

ACCESSION NUMBER:

2004:18737 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

NUMBER KIND DATE
US 2004014039 A1 20040122

PATENT INFORMATION: APPLICATION INFO.:

PATENT ASSIGNEE(S):

US 2002-158057 A1 20020531 (10)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2001-764890, filed on 17

Jan 2001, PENDING

|                       | NUMBER                             | DATE     |      |
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DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

LINE COUNT: 26776

L11 ANSWER 8 OF 221 USPATFULL on STN

Albumin fusion proteins ΤI

The present invention encompasses albumin fusion AB .. proteins. Nucleic acid molecules encoding the albumin fusion proteins of the invention are also encompassed by the invention, as are vectors containing these nucleic acids, host cells transformed with these nucleic acids vectors, and methods of making the albumin fusion proteins of the invention and using these nucleic acids, vectors, and/or host cells. Additionally the present invention encompasses pharmaceutical compositions comprising albumin fusion proteins and methods of treating, preventing, or ameliorating diseases, disordrs or conditions using albumin fusion proteins of the invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13611 USPATFULL Albumin fusion proteins

TITLE: INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Haseltine, William A., Washington, DC, UNITED STATES

NUMBER KIND DATE \_\_\_\_\_ US 2004010134 A1 20040115

PATENT INFORMATION: APPLICATION INFO.: US 2001-833245 A1 20010412 (9)

DATE NUMBER \_\_\_\_\_ PRIORITY INFORMATION: US 2000-256931P 20001221 (60) 20000425 (60) US 2000-199384P US 2000-229358P 20000412 (60)

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 18 Drawing Page(s)

LINE COUNT: 25066

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 9 OF 221 USPATFULL on STN

7 Human ovarian and ovarian cancer associated proteins TI

This invention relates to newly identified ovarian or ovarian cancer AB related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "ovarian cancer antigens",

and the use of such ovarian antigens for detecting disorders of the reproductive system, particularly the presence of ovarian cancer and ovarian cancer metastases. This invention relates to ovarian cancer antigens as well as vectors, host cells, antibodies directed to ovarian cancer antigens and the recombinant methods and synthetic methods for producing the same. Also provided are diagnostic methods for detecting, treating, preventing and/or prognosing disorders related to the ovary, including ovarian cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of ovarian cancer antigens of the invention. The present invention further relates to inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13598 USPATFULL

TITLE:

AB

7 Human ovarian and ovarian cancer associated proteins Birse, Charles E., North Potomac, MD, UNITED STATES

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

|                     | NUMBER         | KIND | DATE     |      |
|---------------------|----------------|------|----------|------|
|                     |                |      |          |      |
| PATENT INFORMATION: | US 2004010121  | A1   | 20040115 |      |
| APPLICATION INFO.:  | US 2003-333900 | A1   | 20030124 | (10) |
|                     | WO 2001-US8585 |      | 20010316 |      |

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM: LINE COUNT: 16023

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 10 OF 221 USPATFULL on STN

Use of bioactive glass compositions to stimulate osteoblast production TI. Compositions comprising bioactive glass compositions or extracts thereof which include ions in an appropriate concentration and ratio that they enhance osteoblast production, and methods of preparation and use thereof, are disclosed. The compositions can be included in implantable devices that are capable of inducing tissue formation in autogeneic, allogeneic and xenogeneic implants, for example as coatings and/or matrix materials. Examples of such devices include prosthetic implants, sutures, stents, screws, plates, tubes, and the like. Aqueous extracts of the bioactive glass compositions, which extracts are capable of stimulating osteoblast production, are also disclosed. The compositions can be used, for example, to induce local tissue formation from a progenitor cell in a mammal, for accelerating allograft repair in a mammal, for promoting in vivo integration of an implantable prosthetic device to enhance the bond strength between the prosthesis and the existing target tissue at the joining site, and for treating tissue degenerative conditions.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:13078 USPATFULL

TITLE:

Use of bioactive glass compositions to stimulate

osteoblast production

INVENTOR (S):

Hench, Larry L, London, UNITED KINGDOM Polak, Julia M, London, UNITED KINGDOM Buttery, Lee D.k., London, UNITED KINGDOM

Xynos, Ioannis D, Nafplion, GREECE

Maroothynaden, Jason, London, UNITED KINGDOM

NUMBER KIND DATE

A1 20040115 A1 20030707 US 2004009598 PATENT INFORMATION:

US 2003-332731 20030707 (10) APPLICATION INFO.:

WO 2001-US21801 20010711

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

BURNS DOANE SWECKER & MATHIS L L P, POST OFFICE BOX LEGAL REPRESENTATIVE:

1404, ALEXANDRIA, VA, 22313-1404

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1 1301 LINE COUNT:

AB

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 11 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ТT

The present invention relates to novel polynucleotides associated with the plasma membrane, the polypeptides encoded by these polynucleotides herein collectively referred to as "plasma membrane associated antigens," and antibodies that immunospecifically bind these polypeptides, and the use of such plasma membrane associated polynucleotides, antigens, and antibodies for detecting, treating, preventing and/or prognosing disorders related to these novel polypeptides. More specifically, isolated nucleic acid molecules are provided encoding novel plasma membrane associated polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing these plasma membrane associated polynucleotides, polypeptides, and/or antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the novel polypeptides of the invention. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The invention further relates to methods and/or compositions for inhibiting or promoting the production and/or function of the polypeptides of the invention.

2004:12971 USPATFULL ACCESSION NUMBER:

Nucleic acids, proteins, and antibodies TITLE:

Birse, Charles E., North Potomac, MD, UNITED STATES INVENTOR (S): Rosen, Craig A., Laytonsville, MD, UNITED STATES

> NUMBER KIND DATE \_\_\_\_\_\_

US 2004009491 A1 20040115 US 2002-264237 A1 20021004 (10) PATENT INFORMATION: APPLICATION INFO.:

Continuation-in-part of Ser. No. WO 2001-US16450, filed RELATED APPLN. INFO.:

on 18 May 2001, PENDING

NUMBER DATE \_\_\_\_\_\_

US 2000-205515P 20000519 (60) PRIORITY INFORMATION:

DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1 LINE COUNT: 18144

L11 ANSWER 12 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies

The present invention relates to novel musculoskeletal system related AΒ polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "musculoskeletal system antigens," and the use of such musculoskeletal system antigens for detecting disorders of

the musculoskeletal system, particularly the presence of cancer and cancer metastases. More specifically, isolated musculoskeletal system associated nucleic acid molecules are provided encoding novel musculoskeletal system associated polypeptides. Novel musculoskeletal system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human musculoskeletal system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the musculoskeletal system, including cancer of musculoskeletal tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:12968 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES Barash, Steven C., Rockville, MD, UNITED STATES

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

KIND NUMBER \_\_\_\_\_ US 2004009488 A1 20040115 US 2002-242515 A1 20020913 (10)

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US 2000-236369P

US 2000-244617P

PATENT INFORMATION: APPLICATION INFO.:

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2001-764877, filed on 17

20000929 (60)

20001101 (60)

Jan 2001, PENDING

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Utility
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DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

24 1

LINE COUNT: 32038 CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 13 OF 221 USPATFULL on STN L11

Methods for the treatment of carcinoma TT

AB The invention concerns compositions and methods for the diagnosis and treatment of neoplastic cell growth and proliferation in mammals, including humans. The invention is based upon the identification of genes that are amplified in the genome of tumor cells, such as renal cell carcinoma. Such gene amplification is expected to be associated with the overexpression of the gene product as compared to normal cells of the same tissue type and contribute to tumorigenesis. Accordingly, the proteins encoded by the amplified genes are believed to be useful targets for the diagnosis and/or treatment (including prevention) of certain cancers, such as renal cell carcinoma, and may act as predictors

of the prognosis of tumor treatment. The present invention is directed to novel methods of diagnosing and treating tumor, such as renal cell carcinoma or Wilms tumor.

ACCESSION NUMBER:

2004:12653 USPATFULL

TITLE:

Methods for the treatment of carcinoma

INVENTOR (S):

Gerritsen, Mary E., San Mateo, CA, UNITED STATES

Peale, Franklin V., JR., San Carlos, CA, UNITED STATES

Wu, Thomas D., San Francisco, CA, UNITED STATES

PATENT ASSIGNEE(S):

GENENTECH, INC. (U.S. corporation)

KIND DATE NUMBER

PATENT INFORMATION:

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APPLICATION INFO.:

US 2004009171 A1 20040115 US 2003-372683 A1 20030221 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2002-271690, filed

on 16 Oct 2002, PENDING

DATE NUMBER

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PRIORITY INFORMATION:

US 2001-344534P 20011018 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

GENENTECH, INC., 1 DNA WAY, SOUTH SAN FRANCISCO, CA,

94080

NUMBER OF CLAIMS:

57

EXEMPLARY CLAIM:

AB

1 6662

LINE COUNT:

L11 ANSWER 14 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies TΙ

The present invention relates to novel ovarian related polynucleotides, the polypeptides encoded by these polynucleotides herein collectively referred to as "ovarian antigens," and antibodies that immunospecifically bind these polypeptides, and the use of such ovarian polynucleotides, antigens, and antibodies for detecting, treating, preventing and/or prognosing disorders of the reproductive system, particularly disorders of the ovaries and/or breast, including, but not limited to, the presence of ovarian and/or breast cancer and ovarian and/or breast cancer metastases. More specifically, isolated ovarian nucleic acid molecules are provided encoding novel ovarian polypeptides. Novel ovarian polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human ovarian polynucleotides, polypeptides, and/or antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the ovaries and/or breast, including ovarian and/or breast cancer, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The invention further relates to methods and/or compositions for inhibiting or promoting the production and/or function of the polypeptides of the invention.

ACCESSION NUMBER:

2004:7345 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR (S):

Birse, Charles E., North Potomac, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES

|        | -            |    | NUMBER     | KIND | DATE         |
|--------|--------------|----|------------|------|--------------|
|        |              |    |            | <br> | <del>-</del> |
| PATENT | INFORMATION: | US | 2004005579 | A1   | 20040108     |

APPLICATION INFO.: US 2002-264049 A1 20021004 (10)

RELATED APPLN. INFO.: Continuation-in-part of Ser. No. WO 2001-US18569, filed

## on 7 Jun 2001, PENDING

NUMBER DATE -----

PRIORITY INFORMATION:

US 2000-209467P 20000607 (60)

DOCUMENT TYPE:

Utility

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

24 1

LINE COUNT:

18130

L11 ANSWER 15 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies ΤI

The present invention relates to novel proteins. More specifically, AΒ isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:7343 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

20001208 (60)

20000927 (60)

PATENT ASSIGNEE(S):

STATES (U.S. corporation)

| NUMBER | KIND | DATE     |  |  |
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PATENT INFORMATION: APPLICATION INFO .:

US 2004005577 A1 20040108

RELATED APPLN. INFO.:

US 2002-242747 A1 20020913 (10) Continuation of Ser. No. US 2001-764881, filed on 17

Jan 2001, PENDING

US 2000-251869P

US 2000-235834P

|          | ·            |    | NUMBER       | DATE     |      |
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Utility
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DOCUMENT TYPE:

FILE SEGMENT:

LEGAL REPRESENTATIVE:

APPLICATION
HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

LINE COUNT: 27694

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 16 OF 221 USPATFULL on STN

Nucleic acids, proteins, and antibodies

TI AB

The present invention relates to novel cardiovascular system related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "cardiovascular system antigens," and the use of such cardiovascular system antigens for detecting disorders of the cardiovascular system, particularly the presence of cancer of cardiovascular system tissues and cancer metastases. More specifically, isolated cardiovascular system associated nucleic acid molecules are provided encoding novel cardiovascular system associated polypeptides. Novel cardiovascular system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human cardiovascular system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the cardiovascular system, including cancer of cardiovascular system tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:7341 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S): Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES, 20850 (U.S. corporation)

PATENT ASSIGNEE(S):

NUMBER KIND DATE

PATENT INFORMATION: APPLICATION INFO.:

US 2004005575 A1 20040108 US 2002-227577 A1 20020826 (10)

RELATED APPLN. INFO.:

Continuation of Ser. No. US 2002-91504, filed on 7 Mar 2002, PENDING Continuation of Ser. No. US 2001-764869,

20000921 (60)

20000921 (60)

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filed on 17 Jan 2001, ABANDONED

|          |              |    | NUMBER       | DATE     | ,    |
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Utility
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DOCUMENT TYPE:

FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24
EXEMPLARY CLAIM: 1
LINE COUNT: 28742

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 17 OF 221 USPATFULL on STN

TI Functional MRI agents for cancer imaging

AB The invention relates to novel magnetic resonance imaging contrast agents for imaging cancer.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:4285 USPATFULL

TITLE:

INVENTOR(S):

Functional MRI agents for cancer imaging Meade, Thomas J., Altadena, CA, United States Fraser, Scott, La Canada, CA, United States

Jacobs, Russell, Arcadia, CA, United States

PATENT ASSIGNEE(S):

Research Corporation Technologies, Inc., Tucson, AZ,

United States (U.S. corporation)

NUMBER KIND DATE \_\_\_\_\_ \_\_\_

PATENT INFORMATION:

US 6673333 B1

20040106

APPLICATION INFO.:

US 2000-715859

20001117 (9)

NUMBER DATE

PRIORITY INFORMATION:

US 2000-201816P 20000504 (60)

DOCUMENT TYPE:

Utility GRANTED

FILE SEGMENT: PRIMARY EXAMINER:

Hartley, Michael G.

LEGAL REPRESENTATIVE:

Dorsey & Whitney LLP, Silva, Robin M., Kossiak, Renee

Μ.

NUMBER OF CLAIMS:

10

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

7 Drawing Figure(s); 5 Drawing Page(s)

LINE COUNT:

2422

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 18 OF 221 USPATFULL on STN

50 human secreted proteins TТ

The present invention relates to novel human secreted proteins and AB isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating diseases, disorders, and/or conditions related to these novel human secreted proteins.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2004:2568 USPATFULL

TITLE:

INVENTOR(S):

50 human secreted proteins Moore, Paul A., Germantown, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES LaFleur, David W., Washington, DC, UNITED STATES Shi, Yanggu, Gaithersburg, MD, UNITED STATES Rosen, Craig A., Laytonsville, MD, UNITED STATES Olsen, Henrik S., Gaithersburg, MD, UNITED STATES Ebner, Reinhard, Gaithersburg, MD, UNITED STATES

PATENT ASSIGNEE(S):

Brewer, Laurie A., St. Paul, MN, UNITED STATES Human Genome Sciences, Inc., Rockville, MD (U.S.

corporation)

KIND DATE NUMBER \_\_\_\_\_\_

PATENT INFORMATION:

APPLICATION INFO.:

US 2004002591 A1 20040101 US 2002-47021 A1 20020117 (10)

RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 2000-722329, filed on 28 Nov 2000, PENDING Continuation of Ser. No. US

1999-262109, filed on 4 Mar 1999, ABANDONED

Continuation-in-part of Ser. No. WO 1998-US18360, filed

on 3 Sep 1998, PENDING

NUMBER DATE PRIORITY INFORMATION: US 2001-262066P 20010118 (60)

19970905 (60) US 1997-57626P

US 1997-57663P 19970905 (60) 19970905 (60)

US 1997-57669P 19970912 (60) US 1997-58666P

US 1997-58667P 19970912 (60)

19970912 (60) US 1997-58973P

19970912 (60) US 1997-58974P US 1998-90112P 19980622 (60)

Utility DOCUMENT TYPE: FILE SEGMENT: APPLICATION

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE, LEGAL REPRESENTATIVE:

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 23 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 2 Drawing Page(s)

33379 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ANSWER 19 OF 221 USPATFULL on STN

Novel human gene relating to respiratory diseases, obesity, and ΤI

inflammatory bowel disease

This invention relates to genes identified from human chromosome AB 20p13-p12, which are associated with various diseases, including asthma. The invention also relates to the nucleotide sequences of these genes, isolated nucleic acids comprising these nucleotide sequences, and isolated polypeptides or peptides encoded thereby. The invention further relates to vectors and host cells comprising the disclosed nucleotide sequences, or fragments thereof, as well as antibodies that bind to the encoded polypeptides or peptides. Also related are ligands that modulate the activity of the disclosed genes or gene products. In addition, the invention relates to methods and compositions employing the disclosed nucleic acids, polypeptides or peptides, antibodies, and/or ligands for use in diagnostics and therapeutics for asthma and other diseases.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2004:2447 USPATFULL

ACCESSION NUMBER:

Novel human gene relating to respiratory diseases, TITLE:

obesity, and inflammatory bowel disease Keith, Tim, Bedford, MA, UNITED STATES

INVENTOR (S): Little, Randall D., Newtonville, MA, UNITED STATES

Eerdewegh, Paul Van, Weston, MA, UNITED STATES

Dupuis, Josee, Newton, MA, UNITED STATES

Del Mastro, Richard G., Norfolk, MA, UNITED STATES

Simon, Jason, Westfield, NJ, UNITED STATES Allen, Kristin, Hopkinton, MA, UNITED STATES Pandit, Sunil, Gaithersburg, MD, UNITED STATES

| NUMBER | KIND | DATE |  |  |  |  |
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PATENT INFORMATION: US 2004002470 A1 20040101

US 2002-277216 A1 20021017 (10) APPLICATION INFO.:

Continuation-in-part of Ser. No. US 2002-126022, filed RELATED APPLN. INFO.: on 19 Apr 2002, PENDING Continuation-in-part of Ser. No. US 2001-834597, filed on 13 Apr 2001, PENDING Continuation-in-part of Ser. No. US 2000-548797, filed

on 13 Apr 2000, PENDING

Utility DOCUMENT TYPE: APPLICATION FILE SEGMENT:

MORGAN & FINNEGAN, L.L.P., 345 PARK AVENUE, NEW YORK, LEGAL REPRESENTATIVE:

NY, 10154

NUMBER OF CLAIMS: 45 EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 162 Drawing Page(s)

LINE COUNT: 15810

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 20 OF 221 USPATFULL on STN

TI Detection and modulation of Slit and roundabount (Robo) mediated

angiogenesis and uses thereof

AB This invention is generally in the field of methods for diagnosis, treatment and prevention of various disorders involving the Slit2

mediated angiogenesis.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:335332 USPATFULL

TITLE:

Detection and modulation of Slit and roundabount (Robo)

mediated angiogenesis and uses thereof

INVENTOR (S):

Geng, Jian-Guo, Portage, MI, UNITED STATES

NUMBER DATE

PRIORITY INFORMATION:

US 2002-362485P 20020308 (60)

DOCUMENT TYPE:

Utility APPLICATION

FILE SEGMENT: LEGAL REPRESENTATIVE:

Peng Chen, Morrison & Foerster LLP, Suite 500, 3811

Valley Centre Drive, San Diego, CA, 92130-2332

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 4 D

4 Drawing Page(s)

LINE COUNT:

AB

1337

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 21 OF 221 USPATFULL on STN

TI Nucleic acids, proteins, and antibodies

The present invention relates to novel excretory system related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "excretory system antigens," and the use of such excretory system antigens for detecting disorders of the excretory system, particularly the presence of cancer of excretory system tissues and cancer metastases. More specifically, isolated excretory system associated nucleic acid molecules are provided encoding novel excretory system associated polypeptides. Novel excretory system polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human excretory system associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the excretory system, including cancer of excretory system tissues, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:334955 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES

PATENT ASSIGNEE(S):

Human Genome Sciences, Inc., Rockville, MD, 20850 (U.S. corporation)

DATE

|  | NUMBER   | KIND          | DATE     |                  |             |
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| PATENT INFORMATION:<br>APPLICATION INFO.:<br>RELATED APPLN. INFO.: | US 2003235831<br>US 2002-242355<br>Continuation of<br>Jan 2001, PENDIN | Al<br>Ser. No | 20020913 | (10)<br>·764897, | filed on 17 |

NUMBER

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DOCUMENT TYPE: Utility FILE SEGMENT: APPLICATION

LEGAL REPRESENTATIVE: HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24
EXEMPLARY CLAIM: 1
LINE COUNT: 22457

AB

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 22 OF 221 USPATFULL on STN

TI Nucleic acids, proteins, and antibodies

The present invention relates to novel proteins. More specifically, isolated nucleic acid molecules are provided encoding novel polypeptides. Novel polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human polynucleotides and/or polypeptides, and antibodies. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to these novel polypeptides. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting or enhancing the production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER: 2003:334953 USPATFULL

TITLE: Nucleic acids, proteins, and antibodies INVENTOR(S): Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES
Rosen, Craig A., Laytonsville, MD, UNITED STATES
Birse, Charles E., North Potomac, MD, UNITED STATES

PATENT ASSIGNEE(S): Human Genome Sciences, Inc., Rockville, MD, UNITED

STATES (U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION: US 2003235829 A1 20031225

APPLICATION INFO.: US 2002-227646 A1 20020826 (10) RELATED APPLN. INFO.: Continuation of Ser. No. US 2001-86067

Continuation of Ser. No. US 2001-860670, filed on 21 May 2001, PENDING Continuation-in-part of Ser. No. WO

2001-US1346, filed on 17 Jan 2001, PENDING

NUMBER DATE

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DOCUMENT TYPE:

FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: 24 EXEMPLARY CLAIM: 1

LINE COUNT: 20415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 23 OF 221 USPATFULL on STN

TI Compositions and methods for systemic inhibition of cartilage degradation

AB Methods and compositions for inhibiting articular cartilage degradation. The compositions preferably include multiple chondroprotective agents, including at least one agent that promotes cartilage anabolic activity and at least one agent that inhibits cartilage catabolism. The compositions may also include one or more pain and inflammation

inhibitory agents. The compositions may be administered systemically, such as to treat patients at risk of cartilage degradation at multiple joints, and suitably may be formulated in a carrier or delivery vehicle that is targeted to the joints. Alternatively the compositions may be injected or infused directly into the joint.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

2003:334713 USPATFULL ACCESSION NUMBER:

Compositions and methods for systemic inhibition of TITLE:

cartilage degradation

Demopulos, Gregory A., Mercer Island, WA, UNITED STATES INVENTOR(S):

Palmer, Pamela Pierce, San Francisco, CA, UNITED STATES

Herz, Jeffrey M., Mill Creek, WA, UNITED STATES

Omeros Corporation (U.S. corporation) PATENT ASSIGNEE(S):

KIND NUMBER DATE ----- -----US 2003235589 A1 20031225 US 2003-356649 A1 20030131 PATENT INFORMATION: APPLICATION INFO.:

(10)

Continuation-in-part of Ser. No. US 2002-31546, filed RELATED APPLN. INFO.: on 18 Jan 2002, PENDING A 371 of International Ser. No.

WO 2000-US19864, filed on 21 Jul 2000, PENDING

Continuation-in-part of Ser. No. US 2001-839633, filed on 20 Apr 2001, PENDING Continuation-in-part of Ser. No. WO 1999-US26330, filed on 5 Nov 1999, PENDING

Continuation-in-part of Ser. No. WO 1999-US24625, filed

on 20 Oct 1999, PENDING

DATE NUMBER \_\_\_\_\_\_ US 2002-353552P 20020201 (60) PRIORITY INFORMATION: US 1999-144904P 19990721 (60) US 1998-107256P 19981105 (60) US 1998-105026P 19981020 (60)

DOCUMENT TYPE: Utility APPLICATION FILE SEGMENT:

OMEROS MEDICAL SYSTEMS, INC., 1420 FIFTH AVENUE, SUITE LEGAL REPRESENTATIVE:

2675, SEATTLE, WA, 98101

NUMBER OF CLAIMS: 155 EXEMPLARY CLAIM:

NUMBER OF DRAWINGS: 9 Drawing Page(s)

LINE COUNT: 6575

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

#### ANSWER 24 OF 221 USPATFULL on STN 1.11

Nucleic acids, proteins, and antibodies ΤI

AB The present invention relates to novel endocrine related polynucleotides and the polypeptides encoded by these polynucleotides herein collectively known as "endocrine antigens," and the use of such endocrine antigens for detecting disorders of the endocrine system, particularly the presence of cancers of the endocrine system and endocrine cancer metastases. More specifically, isolated endocrine associated nucleic acid molecules are provided encoding novel endocrine associated polypeptides. Novel endocrine polypeptides and antibodies that bind to these polypeptides are provided. Also provided are vectors, host cells, and recombinant and synthetic methods for producing human endocrine associated polynucleotides and/or polypeptides. The invention further relates to diagnostic and therapeutic methods useful for diagnosing, treating, preventing and/or prognosing disorders related to the endocrine system, including cancers of the endocrine system, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying agonists and antagonists of polynucleotides and polypeptides of the invention. The present invention further relates to methods and/or compositions for inhibiting the

production and function of the polypeptides of the present invention.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:330759 USPATFULL

TITLE:

Nucleic acids, proteins, and antibodies

INVENTOR(S):

Rosen, Craig A., Laytonsville, MD, UNITED STATES

Ruben, Steven M., Olney, MD, UNITED STATES

Barash, Steven C., Rockville, MD, UNITED STATES Human Genome Sciences, Inc., Rockville, MD (U.S.

corporation)

NUMBER KIND DATE

PATENT INFORMATION:

PATENT ASSIGNEE(S):

US 2003232975 A1 20031218 US 2002-74024 A1 20020214

APPLICATION INFO.: RELATED APPLN. INFO.: (10)

Continuation of Ser. No. US 2001-764895, filed on 17 Jan 2001, ABANDONED

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Utility
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DOCUMENT TYPE: FILE SEGMENT:

APPLICATION

LEGAL REPRESENTATIVE:

HUMAN GENOME SCIENCES INC, 9410 KEY WEST AVENUE,

ROCKVILLE, MD, 20850

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

24

LINE COUNT:

21828

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

L11 ANSWER 25 OF 221 USPATFULL on STN

TI Proteases

The invention provides human proteases (PRTS) and polynucleotides which identify and encode PRTS. The invention also provides expression vectors, host cells, antibodies, agonists, and antagonists. The invention also provides methods for diagnosing, treating, or preventing disorders associated with aberrant expression of PRTS.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

ACCESSION NUMBER:

2003:330138 USPATFULL

TITLE:

Proteases

INVENTOR(S):

Delegeane, Angelo M., Milpitas, CA, UNITED STATES Gandhi, Ameena R., San Francisco, CA, UNITED STATES Hafalia, April J. A., Santa Clara, CA, UNITED STATES Lu, Dyung Aina M., San Jose, CA, UNITED STATES Arvizu, Chandra S., San Jose, CA, UNITED STATES Tribouley, Catherine M., San Francisco, CA, UNITED

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